

# STIC Search Report Biotech-Chem Library

# STIC Database Tracking Number: 124658

TO: Ben Sackey

Location: REM-5B31&5C18

Art Unit: 1626

Friday, June 18, 2004

Case Serial Number: 10/733134

From: Noble Jarrell

**Location: Biotech-Chem Library** 

**Rem 1B71** 

Phone: 272-2556

Noble.jarrell@uspto.gov

Search Notes	TOTAL SECTION AND ADDRESS OF THE PROPERTY OF T	



124658

Access DB#

# SEARCH REQUEST FORM

## Scientific and Technical Information Center

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f more than one search is submit	******	e searches in order of need.  *****************  is specifically as possible the subject matter to be searched.	Salar Control of the Control
include the elected species or structures, key utility of the invention. Define any terms the known. Please attach a copy of the cover sho	words, synonyms, acrony at may have a special me cet, pertinent claims, and	yms, and registry numbers, and combine with the concept of aning. Give examples or relevant citations, authors, etc, if abstract.	
Title of Invention: Cyclopentan	*(ENE) LEPTEDO	c a herotanoic acids & derivatives thereof useful	
Inventors (please provide full names):	obet M. Bu	wk	Made and Address
Earliest Priority Filing Date: 10/2	28/23		CHANGE ACCORDING TO THE
*For Sequence Searches Only* Please include appropriate serial number.	all pertinent information (	parent, child, divisional, or issued patent numbers) along with the	
A compand & 7-130	4-54-dilydroxy	-23-(3x-methoxy-1E-octenyi)-cyclopantyl]- and metrod of nsing Same	
52-heptenamide	, composition	and me	200000000000000000000000000000000000000
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STAFF USE ONLY	Type of Search	Vendors and cost where applicable	
Searcher: Noble Jorrell	NA Sequence (#)	stn 316	
Searcher Phone #:	AA Sequence (#)	Dialog	
Searcher Location:	Structure (#)	Questel/Orbit	
Date Searcher Picked Up:	Bibliographic	Dr.Link	
Date Completed: 61864	Litigation	Lexis/Nexis	
Searcher Prep & Review Time: 20	Fulltext	Sequence Systems	
Clerical Prep Time:	Patent Family	WWW/Internet	
Online Time:	Other	Other (specify)	

PTO-1590 (8-01)

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(FILE 'HOME' ENTERED AT 11:55:26 ON 18 JUN 2004)

FILE 'HCAPLUS' ENTERED AT 11:55:32 ON 18 JUN 2004 E BURK R/AU

L188 E3,E10,E16,E21-22

L2898 ALLERGAN?/CS,PA

Ъ3 6 L1-2 AND HEPTANOIC ACID?/TI

FILE 'STNGUIDE' ENTERED AT 11:58:48 ON 18 JUN 2004

FILE 'REGISTRY' ENTERED AT 11:59:18 ON 18 JUN 2004

FILE 'HCAPLUS' ENTERED AT 11:59:24 ON 18 JUN 2004 TRA L3 1- RN : L4179 TERMS

FILE 'REGISTRY' ENTERED AT 11:59:24 ON 18 JUN 2004  $L_5$ 179 SEA L4

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FILE 'HCAPLUS' ENTERED AT 13:41:46 ON 18 JUN 2004 USE IS SUBJECT TO THE TERMS OF YOUR STN CUSTOMER AGREEMENT. PLEASE SEE "HELP USAGETERMS" FOR DETAILS. COPYRIGHT (C) 2004 AMERICAN CHEMICAL SOCIETY (ACS)

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FILE COVERS 1907 - 18 Jun 2004 VOL 140 ISS 26 FILE LAST UPDATED: 17 Jun 2004 (20040617/ED)

This file contains CAS Registry Numbers for easy and accurate substance identification.

'OBI' IS DEFAULT SEARCH FIELD FOR 'HCAPLUS' FILE

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ANSWER 1 OF 6 HCAPLUS COPYRIGHT 2004 ACS on STN

AN 2003:609847 HCAPLUS

DN 139:128062

Entered STN: 08 Aug 2003

Method of enhancing hair growth using cyclopentane heptanoic acid compounds

INWoodward, David F.; Vandenburgh, Amanda M.

PA

Allergan, Inc., USA
U.S. Pat. Appl. Publ., 11 pp. SO

CODEN: USXXCO

DTPatent

LA English

ICM A61K031-557 IC

```
ICS A61K031-558; A61K007-06
    424070100; 514568000; 514430000; 514277000; 514449000
    1-12 (Pharmacology)
    Section cross-reference(s): 63
FAN.CNT 1
```

```
PATENT NO.
                      KIND DATE
                                           APPLICATION NO.
                                                             DATE
                            _____
                      _ _ _ _
PΙ
     US 2003147823
                                           US 2003-345788
                       Α1
                            20030807
                                                             20030115
     WO 2003066008
                            20030814
                                           WO 2003-US3363
                       A1
                                                             20030203
         W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN,
             CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, ES, FI, GB, GD, GE, GH,
             GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR,
             LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, OM, PH,
             PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, TJ, TM, TN, TR, TT, TZ,
             UA, UG, UZ, VC, VN, YU, ZA, ZM, ZW, AM, AZ, BY, KG, KZ, MD, RU,
             TJ, TM
         RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AT, BE, BG,
             CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IT, LU, MC,
             NL, PT, SE, SI, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW,
             ML, MR, NE, SN, TD, TG
PRAI US 2002-354425P
                     Р
                            20020204
                            20030115
```

US 2003-345788 Α

MARPAT 139:128062 OS

GΙ

$$R^1$$
 $A-B$ 
 $R^2$ 

AΒ Methods and compns. for stimulating the growth of hair are disclosed wherein said compns. include a cyclopentane heptanoic acid, 2-cycloalkyl or arylalkyl compound I (dashed bonds represent single or double bond which can be in the cis or trans configuration; A = alkylene or alkenylene radical; B = cycloalkyl, aryl; Z = 0; X = N(R4)2; R4 = H, lower alkyl, etc.; R1, R2 = O, OH, O(CO)R6; and R6 = C1-20 (un)saturated acyclic hydrocarbon, etc.). Such compns. are used in treating the skin or scalp of a human or non-human animal. Bimatoprost is preferred for this treatment. In a patient treated for glaucoma with bimatoprost, the eyelashes had increased growth.

ST cyclopentane heptanoate compd enhancing hair growth; eyelash growth bimatoprost

Ι

ΙT Drug delivery systems

> (aerosols; cyclopentane heptanoic acid compds. for enhancing hair growth)

ΙT Alopecia

Animal

Hair

Human

Mammalia

Scalp

Skin

```
(cyclopentane heptanoic acid compds. for enhancing hair growth)
ΙT
     Paraffin oils
     Petrolatum
     Wool wax
     RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
        (cyclopentane heptanoic acid compds. for enhancing hair growth)
ΙT
        (eyelash; cyclopentane heptanoic acid compds. for enhancing hair
        growth)
ΙT
     Hair
        (follicle; cyclopentane heptanoic acid compds. for enhancing hair
        growth)
IT
     Hair preparations
        (growth stimulants; cyclopentane heptanoic acid compds. for enhancing
        hair growth)
IT
     Drug delivery systems
        (lotions; cyclopentane heptanoic acid compds. for enhancing hair
        growth)
IT
     Drug delivery systems
        (ointments, creams; cyclopentane heptanoic acid compds. for enhancing
        hair growth)
IT
     Drug delivery systems
        (powders, topical, dusting; cyclopentane heptanoic acid compds. for
        enhancing hair growth)
IT
     Drug delivery systems
        (solns.; cyclopentane heptanoic acid compds. for enhancing hair growth)
IT
     Waxes
     RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
        (spermaceti; cyclopentane heptanoic acid compds. for enhancing hair
        growth)
IT
     Drug delivery systems
        (topical; cyclopentane heptanoic acid compds. for enhancing hair
        growth)
IT
     5763-58-6D, Cyclopentane heptanoic acid, cycloalkyl or arylalkyl compds.
     155206-00-1, Bimatoprost 155206-00-1D, Bimatoprost, acid addition salts
     RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL
     (Biological study); USES (Uses)
        (cyclopentane heptanoic acid compds. for enhancing hair growth)
ΤТ
     57-55-6, Propylene glycol, biological studies 64-17-5, Ethanol,
     biological studies
                          75-71-8, Dichlorodifluoromethane
                                                             99-76-3,
     Methylparaben 872-50-4, N-Methyl pyrrolidone, biological studies
     1314-13-2, Zinc oxide, biological studies
                                                 1320-37-2,
     Dichlorotetrafluoroethane
                               7732-18-5, Water, biological studies
     8011-96-9, Calamine
                          8049-07-8, Tegacid
                                               9005-65-6, Polysorbate 80
     14807-96-6, Talc, biological studies
    RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
        (cyclopentane heptanoic acid compds. for enhancing hair growth)
    ANSWER 2 OF 6 HCAPLUS COPYRIGHT 2004 ACS on STN
L3
     1997:576688 HCAPLUS
AN
DN
     127:243271
ED
     Entered STN: 10 Sep 1997
    Non-acidic cyclopentane heptanoic acid 2-cycloalkyl or
TT
     arylalkyl derivatives as therapeutic agents
     Woodward, David L.; Andrews, Steven W.; Burk, Robert M.; Garst,
IN
    Michael E.
PA
    Allergan, USA
     PCT Int. Appl., 44 pp.
SO
     CODEN: PIXXD2
```

Patent

DT

```
English
LA
     ICM A61K031-557
IC
     1-12 (Pharmacology)
CC
     Section cross-reference(s): 2, 26, 63
FAN.CNT 6
                                      APPLICATION NO. DATE
     PATENT NO.
                     KIND DATE
     -----
PΤ
     WO 9730710
                                          WO 1997-US2269 19970213
                      A1
                            19970828
         W: AU, CA, JP
         RW: AT, BE, CH, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE
     , ca, £
2008819 A
AU 9722721
                            19971118 US 1996-605567 19960222
                       A1 19970910
                                           AU 1997-22721
                                                             19970213
PRAI US 1996-605567
                       Α
     US 1996-605567 A
US 1992-948056 A3
US 1993-154244 B1
US 1995-371339 A2
WO 1997-US2269 W
                            19960222
                            19920921
                            19931118
                            19950111
     WO 1997-US2269
                       W
                            19970213
OS
     MARPAT 127:243271
AΒ
     The present invention provides cyclopentane heptanoic acid 2-cycloalkyl or
     arylalkyl compds., which may be substituted in the 1-position with amino,
     amido, ether, or ester groups, e.g., a 1-OH cyclopentane heptanoic acid
     2-(cycloalkyl or arylalkyl) compound The cyclopentane heptanoic acid
     2-(cycloalkyl or arylalkyl) compds. of the present invention are potent
     ocular hypotensives, and are particularly suitable for the management of
     glaucoma. Moreover, the compds. of the invention are smooth muscle
     relaxants with broad application in e.g. systemic hypertensive and
     pulmonary diseases. Preparation of cyclopentane heptenamide-5-cis-2-(3lpha-
     hydroxy-4-m-chlorophenoxy-1-trans-butenyl)-1,5-dihydroxy,
     [1\alpha, 2\beta, 3\alpha, 5\alpha] is described. The ability of the
     compds. of the invention to lower intraocular pressure was determined
st
     cyclopentane heptanoate deriv prepn therapeutic; glaucoma cyclopentane
     heptanoate deriv
ΙT
     Allergy inhibitors
     Antihypertensives
     Cardiovascular agents
     Drug delivery systems
     Glaucoma (disease)
     Lung, disease
        (cyclopentane heptanoic acid 2-cycloalkyl or arylalkyl non-acidic
        derivs. as therapeutic agents)
IT
     Digestive tract
     Respiratory tract
        (disease; cyclopentane heptanoic acid 2-cycloalkyl or arylalkyl
        non-acidic derivs. as therapeutic agents)
IT
    Reproduction, animal
        (disorder; cyclopentane heptanoic acid 2-cycloalkyl or arylalkyl
        non-acidic derivs. as therapeutic agents)
ΙT
    Muscle relaxants
        (smooth; cyclopentane heptanoic acid 2-cycloalkyl or arylalkyl
       non-acidic derivs. as therapeutic agents)
    40665-92-7, Cloprostenol 40665-92-7D, Cloprostenol, esters
IT
                                                                    40666-16-8,
    Fluprostenol 40666-16-8D, Fluprostenol, esters 54276-17-4 54276-21-0
    56988-09-1 155205-90-6 155205-91-7 155205-99-5 155206-00-1
    155206-01-2
                 155206-02-3 155206-03-4 195503-17-4 195503-18-5
    195503-19-6
                 195503-20-9 195503-21-0 195503-22-1 195503-23-2
    195503-24-3
    RL: BAC (Biological activity or effector, except adverse); BSU (Biological
    study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES
     (Uses)
        (cyclopentane heptanoic acid 2-cycloalkyl or arylalkyl non-acidic
```

```
derivs. as therapeutic agents)
IT
     56687-85-5
                73275-76-0
    RL: RCT (Reactant); RACT (Reactant or reagent)
       (reaction; cyclopentane heptanoic acid 2-cycloalkyl or arylalkyl
       non-acidic derivs. as therapeutic agents, and preparation thereof)
Ь3
    ANSWER 3 OF 6 HCAPLUS COPYRIGHT 2004 ACS on STN
AN
    1995:946793 HCAPLUS
DN
    123:339522
ED
    Entered STN: 29 Nov 1995
ΤI
    Cyclopentane (ene) heptenoic or -heptanoic acid
    derivatives useful as therapeutic agents
IN
    Burk, Robert M.
PA
    Allergan, Inc., USA
SO
    PCT Int. Appl.
    CODEN: PIXXD2
DT
    Patent
LA
    English
IC
    ICM C07C405-00
    ICS A61K031-557
CC
    26-3 (Biomolecules and Their Synthetic Analogs)
    Section cross-reference(s): 2
FAN.CNT 3
    PATENT NO.
                    KIND DATE
                                       APPLICATION NO. DATE
    -----
                                        ______
    WO 9518102
PT
                    A1 19950706
                                       WO 1994-US13984 19941206
        W: AT, AU, BB, BG, BR, BY, CA, CH, CN, CZ, DE, DK, ES, FI, GB, HU,
            JP, KP, KR, KZ, LK, LU, LV, MG, MN, MW, NL, NO, NZ, PL, PT, RO,
            RU, SD, SE, SK, UA, UZ, VN
        RW: AT, BE, CH, DE, DK, ES, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE,
            BF, BJ, CF, CG, CI, CM, GA, GN, ML, MR, NE, SN, TD, TG
    US 5545665
                        19960813
                                       US 1993-174535
                    Α
                                                       19931228
    CA 2180008
                    AA 19950706
                                       CA 1994-2180008 19941206
    AU 9513359
                    A1 19950717
                                       AU 1995-13359
                                                        19941206
    AU 696645
                   B2
                          19980917
    EP 737184
                    A1
                          19961016
                                        EP 1995-904818
                                                        19941206
    EP 737184
                   B1
                          19990428
        R: DE, ES, FR, GB, IT
    JP 09507228
                    T2 19970722
                                        JP 1994-518042
                                                        19941206
    ES 2133720
                    T3 19990916
                                       ES 1995-904818
                                                        19941206
    US 5990138
                     Α
                         19991123
                                       US 1999-225034
                                                       19990104
    US 6303658
                    B1 20011016
                                       US 1999-448082
                                                        19991123
    US 2002002150
                    A1 20020103
                                       US 2001-919318
                                                       20010731
    US 6414022
                    B2 20020702
    US 2002143054
                    A1 20021003
                                       US 2002-87867
                                                       20020228
    US 6716876
                    B2 20040406
PRAI US 1993-174535
                    Α
                         19931228
    WO 1994-US13984 W
                         19941206
    US 1995-445842
                    A3 19950711
    US 1996-740883
                    A3 19961104
    US 1997-861414
                    A3
                        19970521
    US 1998-84805
                    A3 19980526
    US 1999-225034
                   Al 19990104
    US 1999-448082
                    A1
                        19991123
    US 2001-919318
                  A1
                          20010731
OS
    MARPAT 123:339522
GI
```

$$R^{1}$$
 $R^{2}$ 
 $R^{3}$ 
 $R$ 
 $R$ 

Title compds. I [R = hdyrocarbon, heteroatom-substituted hydrocarbon; R1-R3 = OH, etherified OH; X = OH, acyloxy, alkoxy, (un) substituted amino; Y = H2, O] are potent ocular hypotensives, and are particularly suitable for the management of glaucoma. Thus, PGF2 $\alpha$  Me ester was alkylated to give a mixture of Me ethers from which the 11-Me ether was isolated. This compound lowered the intraocular pressure in dogs by 6.2 mm in 0.1% solution

ST prostaglandin F2a ether prepn ocular hypotensive

IT Glaucoma (disease)

(preparation of prostaglandin derivs. as ocular hypotensives)

IT 170753-66-9P 170753-73-8P

RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); RCT (Reactant); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); RACT (Reactant or reagent); USES (Uses)

(preparation of prostaglandin derivs. as ocular hypotensives)

IT73726-97-3P 79743-27-4P 136198-86-2P 170753-65-8P 170753-67-0P 170753-68-1P 170753-69-2P 170753-70-5P 170753-71-6P 170753-72-7P 170753-74-9P 170753-75-0P 170753-76-1P 170753-77-2P 170753-78-3P 170753-79-4P 170753-80-7P 170753-81-8P 170753-82-9P 170753-83-0P 170753-84-1P 170753-85-2P 170753-86-3P 170753-87-4P 170753-88-5P 170753-89-6P 170753-90-9P 170753-91-0P 170753-92-1P 170753-93-2P 170753-94-3P 170753-95-4P 170753-96-5P 170753-97-6P 170753-98-7P RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)

(preparation of prostaglandin derivs. as ocular hypotensives)

IT 40834-99-9P 73726-94-0P 73726-96-2P 170753-99-8P

RL: BYP (Byproduct); PREP (Preparation)

(preparation of prostaglandin derivs. as ocular hypotensives)

IT 551-11-1, Prostaglandin F2 $\alpha$  33854-16-9, Prostaglandin F2 $\alpha$ 

methyl ester 53764-90-2 170754-00-4 170754-01-5 170754-02-6

RL: RCT (Reactant); RACT (Reactant or reagent)

(preparation of prostaglandin derivs. as ocular hypotensives)

IT 63598-54-9P 65844-25-9P 65844-26-0P

RL: RCT (Reactant); SPN (Synthetic preparation); PREP (Preparation); RACT (Reactant or reagent)

(preparation of prostaglandin derivs. as ocular hypotensives)

- L3 ANSWER 4 OF 6 HCAPLUS COPYRIGHT 2004 ACS on STN
- AN 1995:420606 HCAPLUS
- DN 123:983
- ED Entered STN: 17 Mar 1995
- TI 2-Hydrocarbyl sulfonamidomethyl Cyclopentane(ene) heptanoic
  acids and 2-hydrocarbyl sulfonamidomethyl cyclopentane(ene)
  heptenoic acids and their derivatives as therapeutic agents for ocular
  hypotension
- IN Andrews, Steven W.

```
Allergan, Inc., USA
PΑ
SO
     U.S., 13 pp.
     CODEN: USXXAM
DT
     Patent
     English
LA
     ICM A61K031-215
IC
     ICS A61K031-195; C07C069-74; C07C405-00
NCL
     514530000
     1-12 (Pharmacology)
     Section cross-reference(s): 24
FAN.CNT 1
     PATENT NO.
                   KIND DATE
                                            APPLICATION NO. DATE
     ---- ----
                             ----
                                             -----
     US 5387608 A 19950207 US 1993-108209
WO 9505178 A1 19950223 WO 1994-US9206
PΤ
                                                               19930817
                                                               19940816
         W: AT, AU, BB, BG, BR, BY, CA, CH, CN, CZ, DE, DK, ES, FI, GB, HU, JP, KP, KR, KZ, LK, LU, LV, MG, MN, MW, NL, NO, NZ, PL, PT, RO,
             RU, SD, SE, SK, UA, UZ, VN
         RW: AT, BE, CH, DE, DK, ES, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE,
             BF, BJ, CF, CG, CI, CM, GA, GN, ML, MR, NE, SN, TD, TG
                      AA 19950223 CA 1994-2169744 19940816
A1 19950314 AU 1994-75657 19940816
     CA 2169744
     AU 9475657
                  A1
     EP 714303
                           19960605
                                            EP 1994-925885
                                                               19940816
         R: DE, ES, FR, GB, IT
     JP 09502964 T2 19970325 JP 1994-507129 US 5457131 A 19951010 US 1994-292543
                                                               19940816
     US 5457131
                       Α
                             19951010
                                             US 1994-292543
                                                               19940818
PRAI US 1993-108209
                             19930817
     WO 1994-US9206
                             19940816
     MARPAT 123:983
OS
AB
     The title compds. (Markush included) are useful as ocular hypotensives.
     Preparation of the compds. of the invention is described, and intraocular
     pressure-lowering effects of e.g. [1\alpha, 2\beta, 3\alpha, 5\alpha]-5-
     cis-2-(phenylethylsulfonamidomethyl)-3,5-dihydroxycyclopentylheptenoic
     acid are given.
ST
     cyclopentane heptanoate sulfonamidomethyl deriv hypotensive eye;
     cyclopentene heptanoate sulfonamidomethyl deriv hypotensive eye;
     heptanoate cyclopentane sulfonamidomethyl deriv hypotensive eye;
     heptenoate cyclopentane sulfonamidomethyl deriv hypotensive eye
IT
     Glaucoma (disease)
        (hydrocarbyl sulfonamidomethyl cyclopentane(ene) heptanoic acids and
        hydrocarbyl sulfonamidomethyl cyclopentane (ene) heptenoic acids and
        their derivs. as therapeutic agents for ocular hypotension, and their
        preparation)
IT
     Pharmaceutical dosage forms
        (ophthalmic, hydrocarbyl sulfonamidomethyl cyclopentane (ene) heptanoic
        acids and hydrocarbyl sulfonamidomethyl cyclopentane(ene) heptenoic
        acids and their derivs. as therapeutic agents for ocular hypotension,
        and their preparation)
IT
     161834-06-6P
     RL: BAC (Biological activity or effector, except adverse); BSU (Biological
     study, unclassified); RCT (Reactant); SPN (Synthetic preparation); THU
     (Therapeutic use); BIOL (Biological study); PREP (Preparation); RACT
     (Reactant or reagent); USES (Uses)
        (hydrocarbyl sulfonamidomethyl cyclopentane (ene) heptanoic acids and
        hydrocarbyl sulfonamidomethyl cyclopentane(ene) heptenoic acids and
        their derivs. as therapeutic agents for ocular hypotension, and their
        preparation)
IT
     161834-09-9P
     RL: BAC (Biological activity or effector, except adverse); BSU (Biological
     study, unclassified); SPN (Synthetic preparation); THU (Therapeutic use);
```

```
BIOL (Biological study); PREP (Preparation); USES (Uses)
         (hydrocarbyl sulfonamidomethyl cyclopentane(ene) heptanoic acids and
         hydrocarbyl sulfonamidomethyl cyclopentane(ene) heptenoic acids and
         their derivs. as therapeutic agents for ocular hypotension, and their
         preparation)
 IT
      98-09-9, Phenylsulfonyl chloride
                                         98-59-9, Toluenesulfonyl chloride
      124-63-0, Methanesulfonyl chloride 1191-15-7, Diisobutyl aluminum
                1939-99-7, Benzylsulfonyl chloride
                                                     2386-60-9, n-Butanesulfonyl
      chloride
                 4025-71-2, Benzeneethanesulfonyl chloride
                                                             6303-18-0,
      1-Pentanesulfonyl chloride
                                   17814-85-6, (4-Carboxybutyl)triphenylphosphon
      ium bromide
                    63014-04-0, Benzenepropanesulfonyl chloride
                                                                  113566-26-0
      RL: RCT (Reactant); RACT (Reactant or reagent)
         (hydrocarbyl sulfonamidomethyl cyclopentane(ene) heptanoic acids and
        hydrocarbyl sulfonamidomethyl cyclopentane(ene) heptenoic acids and
        their derivs. as therapeutic agents for ocular hypotension, and their
        preparation)
 IT
      58707-52-1P
                   58707-53-2P
                                                161833-84-7P
                                  58707-54-3P
                                                               161833-85-8P
      161833-86-9P
                    161833-87-0P
                                    161833-88-1P
                                                   161833-89-2P
                                                                  161833-90-5P
      161833-91-6P
                     161833-92-7P
                                    161833-93-8P
                                                   162120-43-6P
     RL: RCT (Reactant); SPN (Synthetic preparation); PREP (Preparation); RACT
      (Reactant or reagent)
         (hydrocarbyl sulfonamidomethyl cyclopentane(ene) heptanoic acids and
        hydrocarbyl sulfonamidomethyl cyclopentane(ene) heptenoic acids and
        their derivs. as therapeutic agents for ocular hypotension, and their
        preparation)
IT
     161833-94-9P
                    161833-95-0P
                                   161833-96-1P
                                                   161833-97-2P
                                                                  161833-98-3P
     161833-99-4P
                    161834-00-0P
                                   161834-01-1P
                                                   161834-02-2P
                                                                  161834-03-3P
     161834-04-4P
                    161834-05-5P
                                   161834-07-7P
     RL: RCT (Reactant); SPN (Synthetic preparation); THU (Therapeutic use);
     BIOL (Biological study); PREP (Preparation); RACT (Reactant or reagent);
     USES (Uses)
        (hydrocarbyl sulfonamidomethyl cyclopentane(ene) heptanoic acids and
        hydrocarbyl sulfonamidomethyl cyclopentane(ene) heptenoic acids and
        their derivs. as therapeutic agents for ocular hypotension, and their
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IT
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                    161834-10-2P
                                   161834-11-3P
     RL: SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological
     study); PREP (Preparation); USES (Uses)
        (hydrocarbyl sulfonamidomethyl cyclopentane(ene) heptanoic acids and
        hydrocarbyl sulfonamidomethyl cyclopentane(ene) heptenoic acids and
        their derivs. as therapeutic agents for ocular hypotension, and their
        preparation)
L3
     ANSWER 5 OF 6 HCAPLUS COPYRIGHT 2004 ACS on STN
     1994:426935 HCAPLUS
AN
DN
     121:26935
ED
     Entered STN: 23 Jul 1994
     7-(5-Substituted cyclopentyl) and (5-substituted cyclopentenyl) heptyl
ΤI
     alcohols, heptylamines and heptanoic acid amines, and
     method of lowering intraocular pressure
IN
     Garst, Michael E.; Burk, Robert
     Allergan, Inc., USA
PA
SO
     PCT Int. Appl., 40 pp.
     CODEN: PIXXD2
DT
     Patent
LΑ
    English
IC
     ICM A61K031-557
    ICS C07C405-00
CC
    1-12 (Pharmacology)
    Section cross-reference(s): 26
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FAN.CNT 1
                                           APPLICATION NO.
                                                             DATE
                      KIND DATE
     PATENT NO.
                                            _____
                            _____
                                                             19931020
                                           WO 1993-US10061
                       A1
                            19940428
PΙ
     WO 9408587
         W: AU, CA, CZ, HU, JP, NO, NZ
         RW: AT, BE, CH, DE, DK, ES, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE
                                           US 1992-964223
                                                             19921021
                            19950131
     US 5385945
                       Α
                                           CA 1993-2147502
                                                             19931020
                            19940428
     CA 2147502
                       AA
                                                             19931020
                                           AU 1994-54094
                            19940509
                       A1
     AU 9454094
                            19960627
     AU 669957
                       В2
                                           EP 1993-924387
                                                             19931020
                            19950809
                       Α1
     EP 665751
                            20020102
                       В1
     EP 665751
         R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IE, IT, LI, LU, MC, NL, PT, SE
                                            JP 1993-510375
                                                             19931020
                       T2
                            19960319
     JP 08502495
                                            AT 1993-924387
                                                             19931020
                            20020115
                       Ε
     AT 211386
                                            ES 1993-924387
                                                             19931020
                            20020801
                       T3
     ES 2170076
                                            US 1994-355463
                                                             19941214
                            19960903
     US 5552434
                       Α
                                            US 1995-572437
                                                             19951214
                            19971007
                       Α
     US 5674910
                                            US 1997-899972
                                                             19970724
                            19980630
     US 5773654
                       Α
                            19921021
PRAI US 1992-964223
                       Α
                            19931020
     WO 1993-US10061
                       W
                            19941214
     US 1994-355463
                       A3
                            19951214
     US 1995-572437
                       A3
     MARPAT 121:26935
OS
GΙ
```

HO 
$$CH(CH_2)_3X$$
  $(CH_2)_nMe$   $OR^1$  I

Compds. I [dotted line = bond or absence of bond; wavy lines = bonds in AΒ cis or trans configuration; R1 = H, COR2 (R2 = C1-6 lower alkyl, carbocyclic aryl, heterocyclic aryl, carbocyclic aryl- or heteroaryl-substituted lower alkyl); X = CONR3R4, CH2OH, CH2OR5, CH2OCOR6, CH2NR3R4 (R3, R4 = H, lower alkyl; R5 = C1-6 lower alkyl; R6 = C1-6 lower alkyl, carbocyclic aryl, heterocyclic aryl, or carbocyclic aryl- or heteroaryl-substituted lower alkyl); n = 0-8] are capable of lowering intraocular pressure in the eye of a mammal. Preparation and intraocular pressure lowering effect of e.g.  $7\alpha$ -[2 $\alpha$ -hydroxy-5 $\beta$ -(3α-hydroxy-1-trans-octenyl)-cyclopentyl]-5-cis-heptenol are included.

cyclopentyl heptyl alc deriv ocular hypotensive; heptylamine cyclopentyl STderiv ocular hypotensive; heptanoic acid amine cyclopentyl deriv glaucoma; cyclopentenyl heptyl alc deriv ocular hypotensive

Glaucoma (disease) TΤ

(treatment of, substituted cyclopentyl and substituted cyclopentenyl heptyl alcs., heptylamines and heptanoic acid amines for)

155827-52-4P 155827-50-2P 155827-49-9P 155827-48-8P 155827-47-7P IT 155827-60-4P 155827-53-5P 155827-56-8P

RL: SPN (Synthetic preparation); PREP (Preparation)

(preparation and intraocular pressure lowering activity of)

155827-55-7P 155827-57-9P 64775-37-7P 31753-19-2P 53228-02-7P IT 155827-59-1P 155827-61-5P

```
RL: RCT (Reactant); SPN (Synthetic preparation); PREP (Preparation); RACT
    (Reactant or reagent)
       (preparation and reaction of, in ocular hypotensive preparation)
                  155827-54-6P 155827-58-0P
    155827-51-3P
IT
    RL: SPN (Synthetic preparation); PREP (Preparation)
       (preparation of, for ocular hypotensive preparation)
                                                 124-40-3, N,N-Dimethylamine,
    75-31-0, Isopropylamine, biological studies
IT
                                      15572-56-2, Isopropylamine
               506-59-2 13345-50-1
    reactions
    hydrochloride 69739-34-0
    RL: RCT (Reactant); RACT (Reactant or reagent)
        (reaction of, in ocular hypotensive preparation)
    ANSWER 6 OF 6 HCAPLUS COPYRIGHT 2004 ACS on STN
L3
    1994:315840 HCAPLUS
AN
    120:315840
DN
    Entered STN: 25 Jun 1994
ED
    Nonacidic cyclopentane heptanoic acid 2-cycloalkyl or
TI
     arylalkyl derivatives for smooth muscle relaxants and for treatment of
     glaucoma
     Woodward, David F.; Andrews, Steven W.; Burk, Robert M.; Garst,
IN
     Michael E.
     Allergan, Inc., USA
PΑ
     PCT Int. Appl., 86 pp.
SO
     CODEN: PIXXD2
     Patent
DT
     English
LA
     ICM A61K031-557
IC
     1-12 (Pharmacology)
     Section cross-reference(s): 24
FAN.CNT 6
                                        APPLICATION NO. DATE
                    KIND DATE
     PATENT NO.
                                         _____
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     _____
                                        WO 1993-US8472 19930909
                           19940331
                     A1
     WO 9406433
PΙ
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             KP, KR, KZ, LK, LU, MG, MN, MW, NL, NO, NZ, PL, PT, RO, RU, SD,
             SE, SK, UA, VN
         RW: AT, BE, CH, DE, DK, ES, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE,
             BF, BJ, CF, CG, CI, CM, GA, GN, ML, MR, NE, SN, TD, TG
                                                          19920921
                                         US 1992-948056
                           19941004
     US 5352708
                      Α
                                                          19930909
                                          EP 1993-921435
                           19950705
     EP 660716
                      A1
                           20011128
     EP 660716
                      В1
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                                    JP 1993-508155
                                                          19930909
     JP 08501310 T2
                           19960213
                                                          19930909
                                         AU 1993-48526
                     B2
                           19970313
     AU 676492
                           19940412
                      A1
     AU 9348526
                                         AT 1993-921435 19930909
                      E
                           20011215
     AT 209494
     ES 2166364
                                         ES 1993-921435
                                                          19930909
                      Т3
                           20020416
                                                          19930909
                                          PT 1993-921435
                     {f T}
                           20020531
     PT 660716
                    A
 PRAI US 1992-948056
                            19920921
                            19930909
     WO 1993-US8472
                      W
     MARPAT 120:315840
 OS
     Cyclopentane heptanoic acid, 2-cycloalkyl or arylalkyl derivs.,
 AB
     substituted in the 1-position with halo, Me, hydroxyl, nitro, amino,
     amido, azido, oxime, cyano, thiol, ether or thioether groups, e.g., a 1-OH
      cyclopentane heptanoic acid, 2-(cycloalkyl or arylalkyl) derivs, are
     disclosed (Markush included). The compds. of the invention are potent
      ocular hypotensives, and are particularly suitable for the management of
      glaucoma. Moreover, the compds. of the invention are smooth muscle
      relaxants with broad application in systemic hypertensive and pulmonary
      diseases; smooth muscle relaxants with application in gastrointestinal
```

```
disease, reproduction, fertility, incontinence, shock, etc. Preparation of
    selected compds. is described, as are radioligand binding studies, effect
    on intraocular pressure, effect on smooth muscle contraction, etc.
     cyclopentane heptanoate cycloalkl arylalkyl deriv glaucoma; smooth muscle
ST
     relaxant cyclopentane heptanoate deriv
     Allergy inhibitors
IT
     Cardiovascular agents
        (nonacidic cyclopentane heptanoic acid cycloalkyl and arylalkyl
        derivs.)
     Glaucoma (disease)
ΙT
     Shock
        (treatment of, nonacidic cyclopentane heptanoic acid cycloalkyl and
        arylalkyl derivs. for)
     Digestive tract
IT
     Reproductive tract
     Respiratory tract
        disease, treatment of, nonacidic cyclopentane heptanoic acid
        cycloalkyl and arylalkyl derivs. for)
     Muscle relaxants
IT
        (smooth, nonacidic cyclopentane heptanoic acid cycloalkyl and arylalkyl
        derivs.)
                                              155205-90-6
                                                             155205-91-7
                                155205-89-3
                  155205-88-2
     56988-09-1
IT
                                                              155205-96-2
                                 155205-94-0
                                              155205-95-1
     155205-92-8 155205-93-9
                                                              155206-01-2
                                               155206-00-1
                                 155205-99-5
     155205-97-3 155205-98-4
     155206-02-3
                   155206-03-4
     RL: BIOL (Biological study)
        (for glaucoma treatment and smooth muscle relaxant)
                                155205-89-3P
     38315-47-8P
                   56687-85-5P
IT
     RL: RCT (Reactant); SPN (Synthetic preparation); PREP (Preparation); RACT
     (Reactant or reagent)
        (preparation and reaction of, in nonacidic cyclopentane heptanoic acid
        cycloalkyl/arylalkyl derivative preparation)
                                                                  155206-05-6P
                                                 155205-95-1P
                                  155205-92-8P
                    155205-90-6P
     155205-88-2P
TT
     RL: SPN (Synthetic preparation); PREP (Preparation)
         (preparation of, for nonacidic cyclopentane heptanoic acid
        cycloalkyl/arylalkyl derivative preparation for glaucoma treatment or smooth
        muscle relaxant)
                  54276-21-0
                               155206-04-5
     38344-08-0
IT
     RL: RCT (Reactant); RACT (Reactant or reagent)
         (reaction of, in nonacidic cyclopentane heptanoic acid
         cycloalkyl/arylalkyl derivative preparation)
                   155206-06-7
     155206-02-3
IT
     RL: BIOL (Biological study)
         (receptor binding competition with, nonacidic cyclopentane heptanoic
         acid cycloalkyl and arylalkyl derivs. for glaucoma treatment or smooth
         muscle relaxant in relation to)
                                                                     67508-08-1
                                                       64775-48-0
                                           64775-47-9
                 33854-16-9 38344-08-0
      551-11-1
 IT
                                             155206-08-9
                                                            155206-09-0
                   96752-55-5
                                155206-07-8
      68192-10-9
                                  155322-19-3 155322-20-6
                    155206-12-5
      155206-10-3
      RL: PRP (Properties)
         (smooth muscle stimulant property of)
 IT
      155206-11-4
      RL: BIOL (Biological study)
         (vasorelaxation response with)
 => => b home
 FILE 'HOME' ENTERED AT 13:42:13 ON 18 JUN 2004
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=> b reg FILE 'REGISTRY' ENTERED AT 12:51:08 ON 18 JUN 2004 USE IS SUBJECT TO THE TERMS OF YOUR STN CUSTOMER AGREEMENT. PLEASE SEE "HELP USAGETERMS" FOR DETAILS. COPYRIGHT (C) 2004 American Chemical Society (ACS)

Property values tagged with IC are from the ZIC/VINITI data file provided by InfoChem.

STRUCTURE FILE UPDATES: 16 JUN 2004 HIGHEST RN 694434-66-7 DICTIONARY FILE UPDATES: 16 JUN 2004 HIGHEST RN 694434-66-7

TSCA INFORMATION NOW CURRENT THROUGH JANUARY 6, 2004

Please note that search-term pricing does apply when conducting SmartSELECT searches.

Crossover limits have been increased. See HELP CROSSOVER for details.

Experimental and calculated property data are now available. For more information enter HELP PROP at an arrow prompt in the file or refer to the file summary sheet on the web at: http://www.cas.org/ONLINE/DBSS/registryss.html

#### => d ide 110

L10 ANSWER 1 OF 1 REGISTRY COPYRIGHT 2004 ACS on STN

RN 170753-89-6 REGISTRY

CN Prosta-5,13-dien-1-amide, 9,11-dihydroxy-15-methoxy-, (5Z,9α,11α,13E,15S)- (9CI) (CA INDEX NAME)

OTHER NAMES:

CN AGN 192151

FS STEREOSEARCH

MF C21 H37 N O4

SR CA

LC STN Files: CA, CAPLUS, USPAT2, USPATFULL

DT.CA CAplus document type: Journal; Patent

RL.P Roles from patents: BIOL (Biological study); PREP (Preparation); USES (Uses)

RL.NP Roles from non-patents: BIOL (Biological study); PROC (Process); PRP (Properties)

Absolute stereochemistry.

Double bond geometry as shown.

\*\*PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT\*\*

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3 REFERENCES IN FILE CA (1907 TO DATE)
               3 REFERENCES IN FILE CAPLUS (1907 TO DATE)
=> d his
     (FILE 'HOME' ENTERED AT 11:55:26 ON 18 JUN 2004)
     FILE 'HCAPLUS' ENTERED AT 11:55:32 ON 18 JUN 2004
                E BURK R/AU
             88 E3, E10, E16, E21-22
L1
L2
            898 ALLERGAN?/CS,PA
              6 L1-2 AND HEPTANOIC ACID?/TI
L3
     FILE 'REGISTRY' ENTERED AT 11:59:18 ON 18 JUN 2004
     FILE 'HCAPLUS' ENTERED AT 11:59:24 ON 18 JUN 2004
                TRA L3 1- RN :
                                   179 TERMS
L4
     FILE 'REGISTRY' ENTERED AT 11:59:24 ON 18 JUN 2004
            179 SEA L4
L5
            183 C21H37NO4
Lб
              2 L6 AND L5
L7
^{\text{L8}}
             17 L6 AND NR=1 AND C5/ES
              5 L8 AND "PROSTA-5,13-DIEN-1-AMIDE"
L9
              1 L9 AND "9,11-DIHYDROXY-15-METHOXY"
L10
                SEL RN L10
              0 E1/CRN
L11
     FILE 'HCAPLUS' ENTERED AT 12:45:27 ON 18 JUN 2004
             3 L10
L12
              2 L12 AND L1
L13
              2 L12 AND L2
L14
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L15
L16
              3 L13-14
              1 L12 NOT L14
L17
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L18
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                E BURK R/AU
           100 E11
L21
           1050 ALLERGAN?/CS,PA
L22
L23
            13 L19 AND L21
           14 L19 AND L22
L24
              1 L19 NOT L23
L25
     FILE 'HCAPLUS' ENTERED AT 13:01:00 ON 18 JUN 2004
L26
              1 L15 AND (PY<=2002 OR PRY<=2002 OR AY<=2002)
     FILE 'USPATFULL, USPAT2' ENTERED AT 13:01:12 ON 18 JUN 2004
              1 L25 AND (PY<=2002 OR PRY<=2002 OR AY<=2002)
L27
    FILE 'HCAPLUS' ENTERED AT 13:39:18 ON 18 JUN 2004
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Searched by Noble Jarrell 272-2556

3 L12 OR L18

349 C21H35NO4

FILE 'REGISTRY' ENTERED AT 13:59:53 ON 18 JUN 2004

37 L29 AND NR=1 AND C5/ES

L28

L29

L30

FILE 'HCAPLUS' ENTERED AT 14:28:04 ON 18 JUN 2004 SEL L28 1- RE

2852 E1-68 L31

FILE 'REGISTRY' ENTERED AT 14:30:47 ON 18 JUN 2004

FILE 'HCAPLUS' ENTERED AT 14:30:57 ON 18 JUN 2004 TRA L31 1- RN : 6154 TERMS L32

FILE 'REGISTRY' ENTERED AT 14:32:21 ON 18 JUN 2004

6154 SEA L32 L33 L34

1 L6 AND L33

L35 0 L29 AND L33

FILE 'USPATFULL, USPAT2' ENTERED AT 14:37:12 ON 18 JUN 2004

SEL L19 1- REP : 71 TERMS L36

L37 428 L36

FILE 'REGISTRY' ENTERED AT 14:39:00 ON 18 JUN 2004

FILE 'USPATFULL, USPAT2' ENTERED AT 14:39:08 ON 18 JUN 2004

TRA L37 1- RN : 10251 TERMS L38

FILE 'REGISTRY' ENTERED AT 14:39:27 ON 18 JUN 2004

L39 10232 SEA L38

4 (L6 OR L29) AND L39 L40

FILE 'HCAPLUS' ENTERED AT 14:41:47 ON 18 JUN 2004

FILE 'REGISTRY' ENTERED AT 14:43:02 ON 18 JUN 2004 2 TERMS

SEL L10 1- CHEM : L41

FILE 'HCAPLUS' ENTERED AT 14:43:03 ON 18 JUN 2004 3 S L41 L42

=> b hcap FILE 'HCAPLUS' ENTERED AT 13:39:44 ON 18 JUN 2004 USE IS SUBJECT TO THE TERMS OF YOUR STN CUSTOMER AGREEMENT. PLEASE SEE "HELP USAGETERMS" FOR DETAILS. COPYRIGHT (C) 2004 AMERICAN CHEMICAL SOCIETY (ACS)

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FILE COVERS 1907 - 18 Jun 2004 VOL 140 ISS 26 FILE LAST UPDATED: 17 Jun 2004 (20040617/ED)

This file contains CAS Registry Numbers for easy and accurate substance identification.

'OBI' IS DEFAULT SEARCH FIELD FOR 'HCAPLUS' FILE

### => d all hitstr 128 tot

L28 ANSWER 1 OF 3 HCAPLUS COPYRIGHT 2004 ACS on STN

AN 2001:703779 HCAPLUS

DN 135:251962

ED Entered STN: 26 Sep 2001

TI Combinations of prostaglandins and brimonidine or derivatives for the treatment of glaucoma and alleviation of elevated intraocular pressure

IN Garst, Michael E.

PA Allergan Sales, Inc., USA

SO U.S., 7 pp., Cont.-in-part of U.S. Ser. No. 710,636, abandoned. CODEN: USXXAM

DT Patent

LA English

IC A61K314-15; A61K312-15; A61K031-19

NCL 514392000

CC 1-8 (Pharmacology)

FAN.CNT 2

FAN.CNT Z				
PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
PI US 6294563	B1	20010925	US 1999-440379	19991115
US 2002010202	A1	20020124	US 2001-903954	20010712
PRAI US 1994-330050	B1	19941027		
US 1996-710636	B2	19960918		
US 1999-440379	<b>A</b> 1	19991115		
OS MARPAT 135:251962	2			
GT				

The invention concerns combinations of alpha adrenergic agents such as brimonidine and its derivs. as represented by formula I below wherein each Y is independently selected from the group consisting of N, N-CH3, O, S and C-R1; R1 is hydrogen, lower alkyl or oxo; R2, R3 and R4 are independently selected from the group consisting of hydrogen, halogen, lower alkyl and lower alkenyl; n is an integer from 1 to 3; and a broken line beside a solid line indicates either a single or a double bond with the proviso that when n=1, both bonds from Y to C-R1 cannot be double bonds, and prostaglandins known in the art to cause lowering of intraocular pressure which are useful in compns., methods of treatment and articles of manufacture for the treatment of glaucoma and alleviation of elevated intraocular pressure and providing neuroprotection (no data).

IT Antiglaucoma agents

(combinations of prostaglandins and brimonidine or derivs. for treatment of glaucoma and alleviation of elevated intraocular pressure)

```
Cytoprotective agents
IT
        (neuroprotectants; combinations of prostaglandins and brimonidine or
        derivs. for treatment of glaucoma and alleviation of elevated
        intraocular pressure)
IT
    Adrenoceptor agonists
        (\alpha-; combinations of prostaglandins and brimonidine or derivs.
        for treatment of glaucoma and alleviation of elevated intraocular
        pressure)
     138282-73-2
IT
     RL: BAC (Biological activity or effector, except adverse); BSU (Biological
     study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES
        (S-1033; combinations of prostaglandins and brimonidine or derivs. for
        treatment of glaucoma and alleviation of elevated intraocular pressure)
                                                21562-57-2, TR-4161
                               745-65-3, PGE1
IT
     363-24-6, PGE2
                     551-11-1
     35121-78-9, Prostacyclin 35536-53-9, 11-Deoxy-PGE2
                                                          35700-23-3
     35700-27-7
                 37786-06-4 38315-43-4
                                           38315-47-8
                                                       38344-08-0
                 39746-25-3, 16,16-Dimethyl-PGE2 40665-92-7, Cloprostenol
     39746-23-1
     40666-16-8, Fluprostenol 51705-19-2 52533-44-5, CP-27987
                                                                   53658-98-3,
     11-Deoxy-16,16-dimethyl-PGE2
                                  53764-89-9 53764-90-2
                                                           53764-90-2D,
             54120-61-5, Prostalene 54315-73-0
                                                    54382-24-0
                                                                54382-74-0
                             59567-61-2, K-10134
                                                    59619-81-7, Etiproston
    59122-46-2, Misoprostol
    59685-85-7, HR-466
                         59803-98-4, Brimonidine
                                                  59982-03-5, CS-412
                                                    62524-99-6, Delprostenate
     60325-46-4, Sulprostone
                             61218-31-3, YPG-209
                                                  67110-79-6, Luprostiol
     62559-74-4, ONO-995
                         64318-79-2, Gemeprost
                         69381-94-8, Fenprostalene
     68382-22-9, HR-601
                                                    69648-08-4, TR-4752
                            71116-82-0, Tiaprost
                                                  73121-56-9, RS-84-135
     69900-71-6, RO-221327
                                                       74317-14-9, TR-4367
                74159-84-5
                              74176-31-1, Alfaprostol
     73647-73-1
                                                77287-05-9, Rioprostil
     74397-12-9, ONO-1206
                          76822-56-5, MDL-646
     79360-43-3, Nocloprost 79378-27-1, CL 116069
                                                     81026-63-3, Enisoprost
                           105595-17-3, ZK 110841
    85923-25-7, SQ 27986
                                                    120891-44-3, ZK 118182
                                135273-43-7
                                              155206-00-1 155925-37-4, RO
     130209-82-4, (Latanoprost)
                                                           155925-56-7, ZK
             155925-39-6, S-747260
                                    155925-50-1, UFO-21
             155925-57-8, 13,14-dihydro-ZK 138519
                                                    170552-18-8, 13,14-dihydro
    ZK 118182 170753-89-6
                            361444-55-5
    RL: BAC (Biological activity or effector, except adverse); BSU (Biological
     study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES
     (Uses)
        (combinations of prostaglandins and brimonidine or derivs. for
        treatment of glaucoma and alleviation of elevated intraocular pressure)
             THERE ARE 35 CITED REFERENCES AVAILABLE FOR THIS RECORD
RE.CNT
RE
(1) Anon; WO 8502841 1985 HCAPLUS
(2) Anon; EP 289349 1988 HCAPLUS
(3) Anon; EP 299914 1989 HCAPLUS
(4) Anon; EP 364417 1990 HCAPLUS
(5) Anon; EP 366279 1990 HCAPLUS
(6) Anon; EP 399839 1990 HCAPLUS
(7) Anon; DE 3923797 1991 HCAPLUS
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(9) Anon; EP 544899 1993 HCAPLUS
(10) Anon; WO 9408585 1994 HCAPLUS
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Searched by Noble Jarrell 272-2556

(15) Chan; US 4994274 1991 HCAPLUS (16) Chang; US 5292517 1994 HCAPLUS

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- (34) Yoles; IOVS 1999, V40(1), P65 MEDLINE
- (35) Yuksel; Ophthalmologica 1999, V213(4), P228 HCAPLUS
- IT 170753-89-6

RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(combinations of prostaglandins and brimonidine or derivs. for treatment of glaucoma and alleviation of elevated intraocular pressure)

RN 170753-89-6 HCAPLUS

CN Prosta-5,13-dien-1-amide, 9,11-dihydroxy-15-methoxy-, (5Z,9α,11α,13E,15S)- (9CI) (CA INDEX NAME)

Absolute stereochemistry.

Double bond geometry as shown.

HO
$$SR$$
 $R$ 
 $R$ 
 $E$ 
 $OMe$ 
 $O$ 
 $CH_2)_4$ 
 $Me$ 

- L28 ANSWER 2 OF 3 HCAPLUS COPYRIGHT 2004 ACS on STN
- AN 2000:893929 HCAPLUS
- DN 134:66281
- ED Entered STN: 21 Dec 2000
- TI Synthetic modification of prostaglandin  $F2\alpha$  indicates different structural determinants for binding to the prostaglandin F receptor versus the prostaglandin transporter
- AU Schuster, Victor L.; Itoh, Shigekazu; Andrews, Steven W.; Burk, Robert M.; Chen, June; Kedzie, Karen M.; Gil, Daniel W.; Woodward, David F.
- CS Department of Medicine, Physiology, Albert Einstein College of Medicine, Bronx, NY, USA
- SO Molecular Pharmacology (2000), 58(6), 1511-1516 CODEN: MOPMA3; ISSN: 0026-895X
- PB American Society for Pharmacology and Experimental Therapeutics
- DT Journal
- LA English
- CC 2-2 (Mammalian Hormones)

Searched by Noble Jarrell 272-2556

Several principles governing the binding of E series prostaglandins to EP AΒ receptors have emerged in recent years. The C-1 carboxyl group binds electrostatically to a conserved arginine residue in the seventh transmembrane segment of the receptor. Prostaglandin E analogs involving bioisosteric replacements of the carboxyl group, such as acylsulfonamide, are also active. In addition, structurally similar esters may also exhibit similar affinity, presumably by virtue of hydrogen bonding. Other regions of the substrate mol. appear to bind to other domains of EP receptors, either via hydrophobic interactions or by hydrogen bonding. Less information is available about the structural requirements for substrate binding to FP receptors. Prostanoids also bind to the prostaglandin transporter PGT. In this case, a conserved C-1 carboxyl group is critically important, since C-1 esters exhibit little affinity. Here we examined the binding of chemical diverse  $PGF2\alpha$  structural analogs to the FP receptor and compared these with binding by the PG transporter. PGT recognized a wide range of anionic substituents. In contrast, the carboxylic acid group was essential for optimal binding to the FP receptor, since replacement by larger moieties with a similar pKa, such as acylsulfonamide and tetrazole, substantially decreased binding affinity. Interestingly, insertion of cyclic substituents in the omega chain increased binding to the FP receptor but reduced affinity for PGT, and substitution for the 15-hydroxyl group produced only a modest reduction in FP receptor binding, but eliminated binding by PGT. Because extracellular  $\text{PGF2}\alpha$  may compete for binding between FP receptors and PGT, these findings have implications for designing  $PGF2\alpha$  analogs for treating disease states.

ST prostaglandin F2alpha analog structure receptor transporter binding

IT Prostanoid receptors

RL: BPR (Biological process); BSU (Biological study, unclassified); BIOL (Biological study); PROC (Process)

(FP; synthetic modification of prostaglandin  $F2\alpha$  indicates different structural determinants for binding to prostaglandin F receptor vs. prostaglandin transporter)

IT Structure-activity relationship

(synthetic modification of prostaglandin F2 $\alpha$  indicates different structural determinants for binding to prostaglandin F receptor vs. prostaglandin transporter)

IT Transport proteins

RL: BPR (Biological process); BSU (Biological study, unclassified); BIOL (Biological study); PROC (Process)

(synthetic modification of prostaglandin  $F2\alpha$  indicates different structural determinants for binding to prostaglandin F receptor vs. prostaglandin transporter)

IT 551-11-1, Prostaglandin F2α 13261-27-3, AGN 190910 40834-96-6, AGN 191995 42743-17-9, AGN 191366 52533-67-2, AGN 191365 53764-90-2 55582-75-7, 17-Phenyl PGF2α 64775-48-0, AGN 191088 68192-10-9, AGN 190911 170753-89-6, AGN 192151 315204-32-1, AGN 194394

RL: BPR (Biological process); BSU (Biological study, unclassified); PRP (Properties); BIOL (Biological study); PROC (Process)

(synthetic modification of prostaglandin  $F2\alpha$  indicates different structural determinants for binding to prostaglandin F receptor vs. prostaglandin transporter)

RE.CNT 33 THERE ARE 33 CITED REFERENCES AVAILABLE FOR THIS RECORD RE

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- IT 170753-89-6, AGN 192151

RL: BPR (Biological process); BSU (Biological study, unclassified); PRP (Properties); BIOL (Biological study); PROC (Process)

(synthetic modification of prostaglandin F2α indicates different structural determinants for binding to prostaglandin F receptor vs. prostaglandin transporter)

RN 170753-89-6 HCAPLUS

CN Prosta-5,13-dien-1-amide, 9,11-dihydroxy-15-methoxy-,  $(5Z,9\alpha,11\alpha,13E,15S)$ - (9CI) (CA INDEX NAME)

Absolute stereochemistry.
Double bond geometry as shown.

L28 ANSWER 3 OF 3 HCAPLUS COPYRIGHT 2004 ACS on STN

AN 1995:946793 HCAPLUS

DN 123:339522

ED Entered STN: 29 Nov 1995

TI Cyclopentane (ene) heptenoic or -heptanoic acid derivatives useful as

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therapeutic agents
     Burk, Robert M.
IN
     Allergan, Inc., USA
PA
     PCT Int. Appl.
SO
     CODEN: PIXXD2
DT
     Patent
LA
     English
IC
     ICM C07C405-00
     ICS A61K031-557
     26-3 (Biomolecules and Their Synthetic Analogs)
     Section cross-reference(s): 2
FAN.CNT 3
                                            APPLICATION NO.
     PATENT NO.
                      KIND
                            DATE
                                            ______
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                            19950706
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             RU, SD, SE, SK, UA, UZ, VN
         RW: AT, BE, CH, DE, DK, ES, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE,
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                       Α
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     AU 9513359
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                                            JP 1994-518042
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                       Α
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     US 2002143054
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PRAI US 1993-174535
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                             19991123
     US 2001-919318
                       A1
                             20010731
os
     MARPAT 123:339522
GΙ
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$$\mathbb{R}^{2}$$
 $\mathbb{R}^{3}$ 
 $\mathbb{R}^{3}$ 

AB Title compds. I [R = hdyrocarbon, heteroatom-substituted hydrocarbon;

I

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R1-R3 = OH, etherified OH; X = OH, acyloxy, alkoxy, (un) substituted amino;
    Y = H2, O] are potent ocular hypotensives, and are particularly suitable
     for the management of glaucoma. Thus, PGF2\alpha Me ester was alkylated
     to give a mixture of Me ethers from which the 11-Me ether was isolated.
     This compound lowered the intraocular pressure in dogs by 6.2 mm in 0.1%
     solution
ST
    prostaglandin F2a ether prepn ocular hypotensive
IT
     Glaucoma (disease)
        (preparation of prostaglandin derivs. as ocular hypotensives)
IT
     170753-66-9P
                    170753-73-8P
     RL: BAC (Biological activity or effector, except adverse); BSU (Biological
     study, unclassified); RCT (Reactant); SPN (Synthetic preparation); THU
     (Therapeutic use); BIOL (Biological study); PREP (Preparation); RACT
     (Reactant or reagent); USES (Uses)
        (preparation of prostaglandin derivs. as ocular hypotensives)
                                               170753-65-8P
                                                                170753-67-0P
ידד
     73726-97-3P
                   79743-27-4P
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     170753-68-1P
                    170753-69-2P
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                                                                  170753-72-7P
     170753-74-9P
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     170753-98-7P
    RL: BAC (Biological activity or effector, except adverse); BSU (Biological
     study, unclassified); SPN (Synthetic preparation); THU (Therapeutic use);
     BIOL (Biological study); PREP (Preparation); USES (Uses)
        (preparation of prostaglandin derivs. as ocular hypotensives)
                                                170753-99-8P
IT
     40834-99-9P
                   73726-94-0P
                                 73726-96-2P
     RL: BYP (Byproduct); PREP (Preparation)
        (preparation of prostaglandin derivs. as ocular hypotensives)
IT
     551-11-1, Prostaglandin F2α
                                   33854-16-9, Prostaglandin F2\alpha
                                                              170754-02-6
     methyl ester
                    53764-90-2
                                 170754-00-4
                                                170754-01-5
    RL: RCT (Reactant); RACT (Reactant or reagent)
        (preparation of prostaglandin derivs. as ocular hypotensives)
IT
                   65844-25-9P
                                 65844-26-0P
     63598-54-9P
     RL: RCT (Reactant); SPN (Synthetic preparation); PREP (Preparation); RACT
     (Reactant or reagent)
        (preparation of prostaglandin derivs. as ocular hypotensives)
     170753-89-6P
IT
     RL: BAC (Biological activity or effector, except adverse); BSU (Biological
     study, unclassified); SPN (Synthetic preparation); THU (Therapeutic use);
     BIOL (Biological study); PREP (Preparation); USES (Uses)
        (preparation of prostaglandin derivs. as ocular hypotensives)
RN
     170753-89-6 HCAPLUS
     Prosta-5,13-dien-1-amide, 9,11-dihydroxy-15-methoxy-,
CN
     (5Z, 9\alpha, 11\alpha, 13E, 15S) - (9CI) (CA INDEX NAME)
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Absolute stereochemistry.

Double bond geometry as shown.

=> b uspatall FILE 'USPATFULL' ENTERED AT 13:40:21 ON 18 JUN 2004 CA INDEXING COPYRIGHT (C) 2004 AMERICAN CHEMICAL SOCIETY (ACS)

FILE 'USPAT2' ENTERED AT 13:40:21 ON 18 JUN 2004 CA INDEXING COPYRIGHT (C) 2004 AMERICAN CHEMICAL SOCIETY (ACS)

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L19 ANSWER 1 OF 14 USPATFULL on STN

AN 2002:259478 USPATFULL

TI Cyclopentane (ENE) heptenoic or heptanoic acids and derivatives thereof useful as therapeutic agents

IN Burk, Robert M., Laguna Beach, CA, UNITED STATES

PA ALLERGAN, INC. (U.S. corporation)

PI US 2002143054 A1 20021003 US 6716876 B2 20040406

AI US 2002-87867 A1 20020228 (10)

Continuation of Ser. No. US 2001-919318, filed on 31 Jul 2001, PENDING Continuation of Ser. No. US 1999-448082, filed on 23 Nov 1999, GRANTED, Pat. No. US 6303658 Continuation of Ser. No. US 1999-225034, filed on 4 Jan 1999, GRANTED, Pat. No. US 5990138 Division of Ser. No. US 1998-84805, filed on 26 May 1998, GRANTED, Pat. No. US 5906989 Division of Ser. No. US 1997-861414, filed on 21 May 1997, GRANTED, Pat. No. US 5798378 Division of Ser. No. US 1996-740883, filed on 4 Nov 1996, GRANTED, Pat. No. US 5681848 Division of Ser. No. US 1995-445842, filed on 11 Jul 1995, GRANTED, Pat. No. US 5587391 Division of Ser. No. US 1993-174535, filed on 28 Dec 1993, GRANTED, Pat. No. US 5545665

DT Utility

FS APPLICATION

LREP ROBERT J. BARAN (T2-7H), ALLERGAN, INC., 2525 Dupont Drive, Irvine, CA, 92612

CLMN Number of Claims: 1

ECL Exemplary Claim: 1

DRWN 4 Drawing Page(s)

LN.CNT 1018

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

The invention relates to 7-[5-hydroxy-2-(hydroxyhydrocarbyl or heteroatom-substituted hydroxy hydrocarbyl)-3-hydroxy-cyclopentyl(enyl)] heptanoic or heptenoic acids and derivatives of said acids, wherein one or more of said hydroxy groups are replaced by an ether group. The compounds of the present invention are potent ocular hypotensives, and are particularly suitable for the management of glaucoma.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

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INCL INCLM: 514/530.000
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INCLS: 514/559.000; 514/659.000

NCL NCLM: 514/530.000

NCLS: 514/546.000; 514/568.000; 514/573.000; 514/613.000; 514/715.000

IC [7]

ICM: A61K031-557

## CHEMICAL ABSTRACTS INDEXING COPYRIGHT 2004 ACS on STN

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US	CA 123:339522 " WO CA 133:252211 US						
	CA 133:252211 US CA 131:5147 WO						
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	26-3 (Biomolecules	ond Their Cur	u nthat	ic Analo	oge)		
CC	Section cross-ref		IICIIEC	ic Anaic	Jys)		
am.	prostaglandin F2a		cular	hymoter	ngive		
ST	Glaucoma (disease)	erner brebu o	Cular	nypoter	IIBIVC		
IT	(preparation of	nrostaglandin	deri	ve 20 (	ocular byo	ntensi	ves)
TM	170753-66-9P 170	prostagrandin	ucii	vs. as	ocurar nyp	Jeenbi	. • • • • •
IT	(preparation of		deri	ve 20 (	ocular hym	ntensi	ves)
IT	73726-97-3P 7974	3-27-4P 136	198-8	vs. as ( 6-2D 1	170753-65-	RP 1	70753-67-0P
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	170753-98-7P	733 24 31 1	,0,55	JJ 11	170733 2	0 01	2.0.00
	(preparation of prostaglandin derivs. as ocular hypotensives)						
$_{ m IT}$	40834-99-9P 7372	6-94-0P 737	26-96	-2P 1	70753-99-8	Р	,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,
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IT	551-11-1, Prostagl	endin F2α 3	3854-	16-9. Pi	rostagland	in F2α	
11	methyl ester 537	64-90-2 170	754 - 0	0-4 11	70754-01-5	170	754-02-6
	(preparation of	prostaglandin	deri	vs. as o	ocular hyp	otensi	ves)
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11	(preparation of				ocular hyp	otensi	ves)
IT	170753-89-6P	P = 0 = 0 = 0 = 0 = 0 = 0 = 0 = 0 = 0			11		•
	(preparation of	prostaglandin	deri	vs. as o	ocular hyp	otensi	ves)
RN	170753-89-6 USPATF				11		
CN	Prosta-5,13-dien-1-		ihydr	oxy-15-r	methoxy-,		
	$(5Z, 9\alpha, 11\alpha, 13E, 15)$	s)- (9CI) (C	A IND	EX NAME)	)		
	(2=, = =, ===, ===, ==	, , , , , , , , , , , , , , , , , , , ,					

Absolute stereochemistry.

Double bond geometry as shown.

HO
$$SR$$
 $R$ 
 $R$ 
 $E$ 
 $OMe$ 
 $O$ 
 $CH_2)_4$ 
 $Me$ 

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ANSWER 2 OF 14 USPATFULL on STN
L19
       2002:4171 USPATFULL
AN
       Cyclopentane (ENE) heptenoic or heptanoic acids and derivatives thereof
TТ
       useful as therapeutic agents
       Burk, Robert M., Laguna Beach, CA, UNITED STATES
IN
       ALLERGAN SALES, INC. (U.S. corporation)
PΑ
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       US 2002002150
PΙ
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       US 6414022
                                 20010731 (9)
                           A1
       US 2001-919318
AI
       Continuation of Ser. No. US 1999-448082, filed on 23 Nov 1999, PENDING Continuation of Ser. No. US 1999-225034, filed on 4 Jan 1999, GRANTED, Pat. No. US 5990138 Division of Ser. No. US 1998-84805, filed on 26 May
RLI
       1998, GRANTED, Pat. No. US 5906989 Division of Ser. No. US 1997-861414,
       filed on 21 May 1997, GRANTED, Pat. No. US 5798378 Division of Ser. No.
       US 1996-740883, filed on 4 Nov 1996, GRANTED, Pat. No. US 5681848
       Division of Ser. No. US 1995-445842, filed on 11 Jul 1995, GRANTED, Pat.
       No. US 5587391 Division of Ser. No. US 1993-174535, filed on 28 Dec
       1993, GRANTED, Pat. No. US 5545665
DT
       Utility
       APPLICATION
FS
       ROBERT J. BARAN (T2-7H), ALLERGAN, INC., 2525 Dupont Drive, Irvine, CA,
LREP
       92612
       Number of Claims: 8
CLMN
       Exemplary Claim: 1
ECL
       4 Drawing Page(s)
DRWN
LN.CNT 1120
CAS INDEXING IS AVAILABLE FOR THIS PATENT.
       The invention relates to 7-[5-hydroxy-2-(hydroxyhydrocarbyl or
       heteroatom-substituted hydroxy hydrocarbyl)-3-hydroxycyclopentyl(enyl)]
       heptanoic or heptenoic acids and derivatives of said acids, wherein one
       or more of said hydroxy groups are replaced by an ether group. The
       compounds of the present invention are potent ocular hypotensives, and
        are particularly suitable for the management of glaucoma.
CAS INDEXING IS AVAILABLE FOR THIS PATENT.
        INCLM: 514/134.000
        INCLS: 514/659.000; 514/715.000; 514/712.000
        NCLM: 514/530.000
NCL
       NCLS: 514/546.000; 514/568.000; 514/573.000; 514/613.000; 514/715.000
        [7]
IC
        ICM: A61K031-66
        ICS: A61K031-5575; A61K031-5578
                                  COPYRIGHT 2004 ACS on STN
CHEMICAL ABSTRACTS INDEXING
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	OS CA 123:339522 * WC	9518102	A1	19950706			
	CA 133:252211 US	6124344	A	20000926			
	CA 131:5147 WC	9925358	A1	19990527			
	* CA Indexing for this r	ecord included	f				
CC 26-3 (Biomolecules and Their Synthetic Analogs)							
Section cross-reference(s): 2							
ST prostaglandin F2a ether prepn ocular hypotensive							
	IT Glaucoma (disease)	Glaucoma (disease)					
(preparation of prostaglandin derivs. as ocular hypotensives)							
	TT 170753-66-9P 170	753-73-8P					
	(preparation of	prostaglandin	deri	vs. as oc	ular hypotensives)		
	тт 73726-97-3P 7974	3-27-4P 136	198-8	6-2P 17	0753-65-8P 170753-67-0P		
	170753-68-1P 170	753-69-2P 1°	70753	-70-5P	170753-71-6P 170753-72-7P		

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170753-77-2P
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                     170753-75-0P
                                     170753-76-1P
      170753-74-9P
                     170753-80-7P
                                                     170753-82-9P
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                     170753-94-3P
                                     170753-95-4P
                                                     170753-96-5P
      170753-93-2P
      170753-98-7P
        (preparation of prostaglandin derivs. as ocular hypotensives)
                    73726-94-0P
                                                 170753-99-8P
                                  73726-96-2P
IT
      40834-99-9P
        (preparation of prostaglandin derivs. as ocular hypotensives)
                                     33854-16-9, Prostaglandin F2\alpha
      551-11-1, Prostaglandin F2α
IT
                                                                170754-02-6
                                   170754-00-4
                                                  170754-01-5
      methyl ester
                     53764-90-2
        (preparation of prostaglandin derivs. as ocular hypotensives)
                    65844-25-9P
                                   65844-26-0P
      63598-54-9P
IT
        (preparation of prostaglandin derivs. as ocular hypotensives)
    170753-89-6P
IT
        (preparation of prostaglandin derivs. as ocular hypotensives)
     170753-89-6 USPATFULL
RN
     Prosta-5,13-dien-1-amide, 9,11-dihydroxy-15-methoxy-,
CN
       (5Z, 9\alpha, 11\alpha, 13E, 15S) - (9CI) (CA INDEX NAME)
```

Absolute stereochemistry. Double bond geometry as shown.

```
ANSWER 3 OF 14 USPATFULL on STN
L19
       2001:179150 USPATFULL
AN
       Cyclopentane heptenoic or heptanoic acids and derivatives thereof useful
TI
       as therapeutic agents
       Burk, Robert M., Laguna Beach, CA, United States
IN
       Allergan Sales, Inc., Irvine, CA, United States (U.S. corporation)
PΆ
                               20011016
       US 6303658
                          В1
PΙ
       US 1999-448082
                               19991123 (9)
ΑI
       Continuation of Ser. No. US 1999-225034, filed on 4 Jan 1999, now
RLT
       patented, Pat. No. US 5990138, issued on 23 Nov 1999 Division of Ser.
       No. US 1998-84805, filed on 26 May 1998, now patented, Pat. No. US
       5906989, issued on 25 May 1999 Division of Ser. No. US 1997-861414,
       filed on 21 May 1997, now patented, Pat. No. US 5798378, issued on 25
       Aug 1998 Division of Ser. No. US 1996-740883, filed on 4 Nov 1996, now
       patented, Pat. No. US 5681848, issued on 28 Oct 1997 Division of Ser.
       No. US 1995-445842, filed on 11 Jul 1995, now patented, Pat. No. US
       5587391, issued on 11 Dec 1996 Division of Ser. No. US 1993-174535,
       filed on 28 Dec 1993, now patented, Pat. No. US 5545665, issued on 13
       Aug 1996
DT
       Utility
       GRANTED
FS
       Primary Examiner: Lambkin, Deborah C.
EXNAM
       Baran, Robert J., Voet, Martin A., Fisher, Carlos A.
LREP
       Number of Claims: 9
CLMN
```

```
Exemplary Claim: 1
ECL
      4 Drawing Figure(s); 4 Drawing Page(s)
DRWN
LN.CNT 1135
CAS INDEXING IS AVAILABLE FOR THIS PATENT.
      The invention relates to 7-[5-hydroxy-2-(hydroxyhydrocarbyl or
      heteroatom-substituted hydroxy hydrocarbyl)-3-hydroxy-cyclopentyl(enyl)]
      heptanoic or heptenoic acids and derivatives of said acids, wherein one
       or more of said hydroxy groups are replaced by an ether group. The
       compounds of the present invention are potent ocular hypotensives, and
       are particularly suitable for the management of glaucoma.
CAS INDEXING IS AVAILABLE FOR THIS PATENT.
       INCLM: 514/613.000
INCL
       INCLS: 564/189.000
              514/613.000
NCL
      NCLM:
       NCLS: 564/189.000
       [7]
TC
       ICM: A01N037-18
EXF
       564/189; 514/613
ARTU
       166
                             COPYRIGHT 2004 ACS on STN
CHEMICAL ABSTRACTS INDEXING
                                      KIND
                                             DATE
                         PATENT
                            9518102 A1 19950706
      CA 123:339522 * WO
OS
                              6124344 A
                                           20000926
                     US
      CA 133:252211
      CA 131:5147 WO
                              9925358 Al 19990527
* CA Indexing for this record included
      26-3 (Biomolecules and Their Synthetic Analogs)
CC
       Section cross-reference(s): 2
      prostaglandin F2a ether prepn ocular hypotensive
ST
      Glaucoma (disease)
TΤ
        (preparation of prostaglandin derivs. as ocular hypotensives)
      170753-66-9P
                     170753-73-8P
IT
        (preparation of prostaglandin derivs. as ocular hypotensives)
                    79743-27-4P 136198-86-2P 170753-65-8P 170753-67-0P
тт
      73726-97-3P
                     170753-69-2P 170753-70-5P 170753-71-6P 170753-72-7P
      170753-68-1P
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                                                                 170753-78-3P
                                   170753-76-1P
                     170753-75-0P
      170753-74-9P
                                   170753-81-8P 170753-82-9P
                                                                  170753-83-0P
                     170753-80-7P
      170753-79-4P
                                    170753-86-3P
                                                   170753-87-4P
                                                                  170753-88-5P
      170753-84-1P
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                                                   170753-96-5P
                                                                  170753-97-6P
      170753-93-2P
      170753-98-7P
        (preparation of prostaglandin derivs. as ocular hypotensives)
                    73726-94-0P 73726-96-2P 170753-99-8P
IT
      40834-99-9P
        (preparation of prostaglandin derivs. as ocular hypotensives)
      551-11-1, Prostaglandin F2\alpha 33854-16-9, Prostaglandin F2\alpha
TТ
                     53764-90-2 170754-00-4 170754-01-5
      methyl ester
        (preparation of prostaglandin derivs. as ocular hypotensives)
      63598-54-9P
                    65844-25-9P
                                  65844-26-0P
IT
        (preparation of prostaglandin derivs. as ocular hypotensives)
    170753-89-6P
TΤ
         (preparation of prostaglandin derivs. as ocular hypotensives)
     170753-89-6 USPATFULL
RN
     Prosta-5,13-dien-1-amide, 9,11-dihydroxy-15-methoxy-,
CN
       (5Z, 9\alpha, 11\alpha, 13E, 15S) - (9CI) (CA INDEX NAME)
       Absolute stereochemistry.
       Double bond geometry as shown.
```

HO 
$$\frac{Z}{(CH_2)_3}$$
  $NH_2$ 
 $R$   $R$   $E$   $S$   $OMe$   $O$   $(CH_2)_4$   $Me$ 

L19 ANSWER 4 OF 14 USPATFULL on STN

AN 2001:163229 USPATFULL

TI Combinations of prostaglandins and brimonidine or derivatives thereof

IN Garst, Michael E., Newport Beach, CA, United States

PA Allergan Sales, Inc., Irvine, CA, United States (U.S. corporation)

PI US 6294563 B1 20010925

AI US 1999-440379 19991115 (9)

RLI Continuation-in-part of Ser. No. US 1998-710636, filed on 17 Mar 1998, now abandoned Continuation of Ser. No. US 1994-330050, filed on 27 Oct 1994, now abandoned

DT Utility FS GRANTED

EXNAM Primary Examiner: Fay, Zohreh

LREP O'Donohue, Cynthia, Fisher, Carlos, Baran, Robert

CLMN Number of Claims: 7 ECL Exemplary Claim: 1

DRWN No Drawings

LN.CNT 605

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

AB The invention concerns combinations of alpha adrenergic agents such as brimonidine and its derivatives as represented by formula (I) below ##STR1##

wherein each Y is independently selected from the group consisting of N, N--CH3, O, S and C--R.sub.1; R.sub.1 is hydrogen, lower alkyl or oxo; R.sub.2, R.sub.3 and R.sub.4 are independently selected from the group consisting of hydrogen, halogen, lower alkyl and lower alkenyl; n is an integer from 1 to 3; and a broken line beside a solid line indicates either a single or a double bond with the proviso that when n=1, both bonds from Y to C--R1 cannot be double bonds,

and prostaglandins known in the art to cause lowering of intraocular pressure

which are useful in compositions, methods of treatment and articles of manufacture for the treatment of glaucoma and alleviation of elevated intraocular pressure and providing neuroprotection.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

INCL INCLM: 514/392.000

INCLS: 514/530.000; 514/573.000; 514/912.000; 514/913.000

NCL NCLM: 514/392.000

NCLS: 514/530.000; 514/573.000; 514/912.000; 514/913.000

IC [7]

ICM: A61K031-415

ICS: A61K031-215; A61K031-19

EXF 514/530; 514/573; 514/912; 514/913; 514/393 ARTU 164

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CHEMICAL ABSTRACTS INDEXING COPYRIGHT 2004 ACS on STN
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                     WO
* US
      CA 125:67796
os
                               6294563 B1 20010925
      CA 135:251962 * US
* CA Indexing for this record included
      1-8 (Pharmacology)
CC
      prostaglandin brimonidine glaucoma intraocular pressure inhibition
ST
      Antiglaucoma agents
IT
        (combinations of prostaglandins and brimonidine or derivs. for
        treatment of glaucoma and alleviation of elevated intraocular pressure)
      Cytoprotective agents
IT
        (neuroprotectants; combinations of prostaglandins and brimonidine or
        derivs. for treatment of glaucoma and alleviation of elevated
        intraocular pressure)
      Adrenoceptor agonists
IT
        (\alpha-; combinations of prostaglandins and brimonidine or derivs.
        for treatment of glaucoma and alleviation of elevated intraocular
        pressure)
      138282-73-2
IT
         (S-1033; combinations of prostaglandins and brimonidine or derivs. for
        treatment of glaucoma and alleviation of elevated intraocular pressure)
                      551-11-1 745-65-3, PGE1 21562-57-2, TR-4161
      363-24-6, PGE2
IT
      35121-78-9, Prostacyclin 35536-53-9, 11-Deoxy-PGE2 35700-23-3 35700-27-7 37786-06-4 38315-43-4 38315-47-8 38344-08-0
      39746-23-1 39746-25-3, 16,16-Dimethyl-PGE2 40665-92-7, Cloprostenol
      40666-16-8, Fluprostenol 51705-19-2 52533-44-5, CP-27987
      53658-98-3, 11-Deoxy-16,16-dimethyl-PGE2 53764-89-9 53764-90-2
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      54382-74-0 59122-46-2, Misoprostol 59567-61-2, K-10134 59619-81-7,
               ton 59685-85-7, HR-466 59803-98-4, Brimonidine 59982-03-5, 60325-46-4, Sulprostone 61218-31-3, YPG-209 62524-99-6,
      Etiproston
      CS-412
      Delprostenate 62559-74-4, ONO-995 64318-79-2, Gemeprost 67110-79-6,
      Luprostiol 68382-22-9, HR-601 69381-94-8, Fenprostalene TR-4752 69900-71-6, RO-221327 71116-82-0, Tiaprost 731
                                                                 73121-56-9,
      RS-84-135 73647-73-1 74159-84-5 74176-31-1, Alfaprostol
       74317-14-9, TR-4367 74397-12-9, ONO-1206 76822-56-5, MDL-646
       77287-05-9, Rioprostil 79360-43-3, Nocloprost 79378-27-1, CL 116069
       81026-63-3, Enisoprost 85923-25-7, SQ 27986 105595-17-3, ZK 110841
       120891-44-3, ZK 118182 130209-82-4, (Latanoprost) 135273-43-7 155206-00-1 155925-37-4, RO 229648 155925-39-6, S-747260
       155925-50-1, UFO-21 155925-56-7, ZK 138519 155925-57-8,
       13,14-dihydro-ZK 138519 170552-18-8, 13,14-dihydro ZK 118182
                    361444-55-5
         (combinations of prostaglandins and brimonidine or derivs. for
         treatment of glaucoma and alleviation of elevated intraocular pressure)
    170753-89-6
         (combinations of prostaglandins and brimonidine or derivs. for
         treatment of glaucoma and alleviation of elevated intraocular pressure)
      170753-89-6 USPATFULL
RN
      Prosta-5,13-dien-1-amide, 9,11-dihydroxy-15-methoxy-,
CN
        (5Z, 9\alpha, 11\alpha, 13E, 15S) - (9CI) (CA INDEX NAME)
```

Absolute stereochemistry.
Double bond geometry as shown.

HO
$$S$$
 $R$ 
 $E$ 
 $O$ 
 $CH_2)_4$ 
 $Me$ 
 $Me$ 

ANSWER 5 OF 14 USPATFULL on STN L19

2001:93539 USPATFULL AN

Cyclopentane heptan(ene)oic acid, 2-heteroarylalk(en)yl derivatives as ΤI therapeutic agents

Burk, Robert M., Laguna Beach, CA, United States IN

Allergan Sales, Inc., United States (U.S. corporation) PA

US 6248773 В1 PΙ

20000822 (9)

20010619

US 2000-643330 AΙ Continuation of Ser. No. US 1999-243344, filed on 1 Feb 1999 RLI Continuation of Ser. No. US 1997-974067, filed on 19 Nov 1997, now patented, Pat. No. US 6124344 Continuation-in-part of Ser. No. US 1997-861414, filed on 21 May 1997, now patented, Pat. No. US 5798378 Division of Ser. No. US 1996-740883, filed on 4 Nov 1996, now patented, Pat. No. US 5681848 Division of Ser. No. US 1995-445842, filed on 11 Jul 1995, now patented, Pat. No. US 5587391 Division of Ser. No. US 1993-174535, filed on 28 Dec 1993, now patented, Pat. No. US 5545665

DT Utility GRANTED FS

Primary Examiner: Seaman, D. Margaret EXNAM

Baran, Robert J., Voet, Martin A., Fisher, Carlos A. LREP

Number of Claims: 28 CLMN Exemplary Claim: 1 ECL

5 Drawing Figure(s); 3 Drawing Page(s) DRWN

LN.CNT 949

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

The invention relates to the use of cyclopentane heptan(ene)oic acid, AB 2-heteroarylalk(en)yl derivatives as ocular hypotensives. The compounds used in accordance with the invention are represented by the following formula I: ##STR1##

wherein the hatched segments represent  $\alpha$  bonds, the solid triangle represents a  $\beta$  bond, wavy line attachments indicate either the alpha  $(\alpha)$  or beta  $(\beta)$  configuration; dashed bonds represent a double bond or a single bond, R is a substituted hetero aryl radical having at least two pendant substituents selected from the group consisting of C.sub.1 to C.sub.6 alkyl; halogen; trifluoromethyl; COR.sup.1; COCF.sub.3; SO.sub.2 NR.sup.1; NO.sub.2 and CN or at least one cyano group; R.sup.1 is hydrogen or a lower alkyl radical having up to six carbon atoms; X is selected from the group consisting of --OR.sup.1 and --N(R.sup.1).sub.2 ; Y is .dbd.0 or represents 2 hydrogen radicals, and the 9, 11, or 15 alkyl esters thereof; provided, however, when said heteroaryl radical is a dichlorothienyl radical, said compound is not a 1-carboxylic acid or amide thereof.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

INCLM: 514/438.000 INCL

INCLS: 514/445.000; 514/448.000; 514/471.000; 514/472.000; 514/473.000;

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514/461.000; 549/061.000; 549/062.000; 549/064.000; 549/066.000;
              549/068.000; 549/070.000; 549/073.000; 549/078.000; 549/474.000;
              549/475.000; 549/476.000; 549/479.000; 549/480.000; 549/483.000;
              549/502.000
             514/438.000
NCL
      NCLM:
             514/445.000; 514/448.000; 514/461.000; 514/471.000; 514/472.000;
      NCLS:
              514/473.000; 549/061.000; 549/062.000; 549/064.000; 549/066.000;
              549/068.000; 549/070.000; 549/073.000; 549/078.000; 549/474.000;
              549/475.000; 549/476.000; 549/479.000; 549/480.000; 549/483.000;
              549/502.000
IC
       [7]
       ICM: A61K031-38
       ICS: A61K031-34; C07D333-38; C07D307-02; C07D333-16
       549/61; 549/62; 549/64; 549/66; 549/68; 549/70; 549/73; 549/78; 549/474;
EXF
       549/475; 549/476; 549/479; 549/480; 549/483; 549/502; 514/445; 514/448;
       514/438; 514/471; 514/472; 514/473; 514/461
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CHEMICAL ABSTRACTS INDEXING
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                            9518102 A1 19950706
      CA 123:339522 * WO
OS
                              6124344 A
      CA 133:252211 US
CA 131:5147 WO
                                           20000926
                             9925358 A1 19990527
* CA Indexing for this record included
      26-3 (Biomolecules and Their Synthetic Analogs)
CC
       Section cross-reference(s): 2
      prostaglandin F2a ether prepn ocular hypotensive
ST
      Glaucoma (disease)
IT
        (preparation of prostaglandin derivs. as ocular hypotensives)
                     170753-73-8P
      170753-66-9P
IT
        (preparation of prostaglandin derivs. as ocular hypotensives)
                                                 170753-65-8P 170753-67-0P
                    79743-27-4P 136198-86-2P
      73726-97-3P
IT
                                                                 170753-72-7P
                                    170753-70-5P
                                                   170753-71-6P
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                                                                 170753-97-6P
                     170753-94-3P
                                    170753-95-4P
      170753-93-2P
      170753-98-7P
        (preparation of prostaglandin derivs. as ocular hypotensives)
                    73726-94-0P 73726-96-2P 170753-99-8P
      40834-99-9P
IT
        (preparation of prostaglandin derivs. as ocular hypotensives)
      551-11-1, Prostaglandin F2\alpha 33854-16-9, Prostaglandin F2\alpha
IT
                     53764-90-2 170754-00-4 170754-01-5
                                                              170754-02-6
      methyl ester
        (preparation of prostaglandin derivs. as ocular hypotensives)
      63598-54-9P
                   65844-25-9P
                                  65844-26-0P
IT
        (preparation of prostaglandin derivs. as ocular hypotensives)
    170753-89-6P
IT
        (preparation of prostaglandin derivs. as ocular hypotensives)
     170753-89-6 USPATFULL
RN
     Prosta-5,13-dien-1-amide, 9,11-dihydroxy-15-methoxy-,
CN
       (5Z, 9\alpha, 11\alpha, 13E, 15S) - (9CI) (CA INDEX NAME)
       Absolute stereochemistry.
```

Double bond geometry as shown.

```
ANSWER 6 OF 14 USPATFULL on STN
L19
       2000:168176 USPATFULL
ΑN
       Cyclopentane heptan(ene)oic acid, 2-heteroarylalkenyl derivatives as
TI
       therapeutic agents
      Burk, Robert M., Laguna Beach, CA, United States
IN
      Allergan Sales, Inc., Irvine, CA, United States (U.S. corporation)
PA
                               20001212
      US 6160129
PΙ
                               19990201 (9)
ΑI
      US 1999-243344
       Continuation of Ser. No. US 1997-974067, filed on 19 Nov 1997 which is a
RLI
       continuation-in-part of Ser. No. US 1997-861414, filed on 21 May 1997,
      now patented, Pat. No. US 5798378 which is a division of Ser. No. US
       1996-740883, filed on 4 Nov 1996, now patented, Pat. No. US 5681848
       which is a division of Ser. No. US 1995-445842, filed on 11 Jul 1995,
       now patented, Pat. No. US 5587391 which is a division of Ser. No. US
       1993-174535, filed on 28 Dec 1993, now patented, Pat. No. US 5545665
       Utility
DT
FS
       Granted
       Primary Examiner: Seaman, D. Margaret
EXNAM
       Baran, Robert J., Fisher, Carlos A., Voet, Martin A.
LREP
       Number of Claims: 10
CLMN
       Exemplary Claim: 1
ECL
       5 Drawing Figure(s); 3 Drawing Page(s)
DRWN
LN.CNT 831
CAS INDEXING IS AVAILABLE FOR THIS PATENT.
       The invention relates to the use of derivatives of F-type prostaglandins
AB
```

as ocular hypotensives. The compounds used in accordance with the invention are represented by the following formula I: ##STR1## wherein wavy line attachments indicate either the alpha  $(\alpha)$  or beta  $(\beta)$  configuration; hatched segments indicate  $\alpha$  configuration; the solid triangle is used to indicate β configuration; dashed bonds represent a double bond, or a single bond; R is a substituted heteroaryl radical having at least two pendant substituents selected from the group consisting of C.sub.1 to C.sub.6 alkyl; halogen; trifluoromethyl; COR.sup.1; COCF.sub.3; SO.sub.2 NR.sup.1; NO.sub.2 and CN or at least one cyano group; R.sup.1 is hydrogen or a lower alkyl radical having up to six carbon atoms, X is selected from the group consisting of --OR.sup.1 and --N(R.sup.1).sub.2; Y is .dbd.0 or represents 2 hydrogen radicals and the 9, 11 or 15 lower alkyl esters thereof; provided, however, when said heteroaryl radical is a dichlorothienyl radical, the compound is not a 1-carboxylic acid or amide thereof. Certain of the compounds represented by Formula I are novel and comprise another aspect of the present invention.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

INCL INCLM: 549/061.000

INCLS: 549/077.000; 549/078.000; 549/079.000; 549/474.000; 549/491.000;

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549/496.000; 549/498.000; 549/502.000; 514/438.000; 514/461.000
NCL
             549/061.000
      NCLM:
            549/077.000; 549/078.000; 549/079.000; 549/474.000; 549/491.000;
      NCLS:
             549/496.000; 549/498.000; 549/502.000
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       [7]
       ICM: A61K031-34
       ICS: A61K031-38; C07D307-02; C07D333-24; C07D333-38
       514/438; 514/461; 546/61; 546/77; 546/78; 546/79; 546/474; 546/491;
EXF
       546/496; 546/498; 546/502
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                    US
                             6124344 A
      CA 133:252211
                             9925358 Al 19990527
      CA 131:5147
                    WO
* CA Indexing for this record included
      26-3 (Biomolecules and Their Synthetic Analogs)
CC
       Section cross-reference(s): 2
      prostaglandin F2a ether prepn ocular hypotensive
st
      Glaucoma (disease)
IT
        (preparation of prostaglandin derivs. as ocular hypotensives)
                     170753-73-8P
      170753-66-9P
ΤТ
        (preparation of prostaglandin derivs. as ocular hypotensives)
                                               170753-65-8P 170753-67-0P
                                136198-86-2P
      73726-97-3P
                   79743-27-4P
TТ
                                                   170753-71-6P 170753-72-7P
                                  170753-70-5P
                    170753-69-2P
      170753-68-1P
                                                   170753-77-2P
                                                                 170753-78-3P
      170753-74-9P
                    170753-75-0P
                                  170753-76-1P
                                                                 170753-83-0P
                                                   170753-82-9P
                    170753-80-7P
                                  170753-81-8P
      170753-79-4P
                                                                 170753-88-5P
                                  170753-86-3P
                                                   170753-87-4P
                   170753-85-2P
      170753-84-1P
                                                   170753-92-1P
                   170753-90-9P
                                    170753-91-0P
      170753-89-6P
                                                                 170753-97-6P
                                                   170753-96-5P
                                   170753-95-4P
      170753-93-2P
                     170753-94-3P
      170753-98-7P
        (preparation of prostaglandin derivs. as ocular hypotensives)
                                73726-96-2P 170753-99-8P
                    73726-94-0P
      40834-99-9P
IT
        (preparation of prostaglandin derivs. as ocular hypotensives)
      551-11-1, Prostaglandin F2\alpha 33854-16-9, Prostaglandin F2\alpha
IT
                                                              170754-02-6
                                  170754-00-4
                                                170754-01-5
      methyl ester
                     53764-90-2
        (preparation of prostaglandin derivs. as ocular hypotensives)
                    65844-25-9P
                                  65844-26-0P
      63598-54-9P
IT
        (preparation of prostaglandin derivs. as ocular hypotensives)
IT
    170753-89-6P
        (preparation of prostaglandin derivs. as ocular hypotensives)
     170753-89-6 USPATFULL
RN
     Prosta-5,13-dien-1-amide, 9,11-dihydroxy-15-methoxy-,
CN
       (5Z, 9\alpha, 11\alpha, 13E, 15S) - (9CI) (CA INDEX NAME)
```

Absolute stereochemistry.

Double bond geometry as shown.

HO
$$S$$
 $R$ 
 $R$ 
 $E$ 
 $O$ 
 $CH_2)_4$ 
 $Me$ 
 $Me$ 

CHEMICAL ABSTRACTS INDEXING

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ANSWER 7 OF 14 USPATFULL on STN
L19
       1999:151241 USPATFULL
AN
       Cyclopentane (ene) heptenoic or heptanoic acids and derivatives thereof
TI
       useful as therapeutic agents
       Burk, Robert M., Laguna Beach, CA, United States
IN
       Allergan, Irvine, CA, United States (U.S. corporation)
PA
                               19991123
PΙ
       US 5990138
                               19990104 (9)
       US 1999-225034
ΑI
       Division of Ser. No. US 1998-84805, filed on 26 May 1998, now patented,
RLI
       Pat. No. US 5906989 which is a division of Ser. No. US 1997-861414,
       filed on 21 May 1997, now patented, Pat. No. US 5798378 which is a
       division of Ser. No. US 1996-740883, filed on 4 Nov 1996, now patented,
       Pat. No. US 5681848 which is a division of Ser. No. US 1995-445842,
       filed on 11 Jul 1995, now patented, Pat. No. US 5587391 which is a
       division of Ser. No. US 1993-174535, filed on 28 Dec 1993, now patented,
       Pat. No. US 5545665
DT
       Utility
FS
       Granted
EXNAM Primary Examiner: Lambkin, Deborah C.
       Number of Claims: 15
       Exemplary Claim: 1
ECL
       4 Drawing Figure(s); 4 Drawing Page(s)
DRWN
LN.CNT 1069
CAS INDEXING IS AVAILABLE FOR THIS PATENT.
       The invention relates to 7-[5-hydroxy-2-(hydroxyhydrocarbyl or
AΒ
       heteroatom-substituted hydroxy hydrocarbyl)-3-hydroxycyclopentyl(enyl)]
       heptanoic or heptenoic acids and derivatives of said acids, wherein one
       or more of said hydroxy groups are replaced by an ether group. The
       compounds of the present invention are potent ocular hypotensives, and
       are particularly suitable for the management of glaucoma.
CAS INDEXING IS AVAILABLE FOR THIS PATENT.
       INCLM: 514/357.000
INCL
       INCLS: 560/121.000
NCL
       NCLM: 514/357.000
       NCLS: 560/121.000
IC
       [6]
       ICM: A61K031-215
       ICS: C07C069-74
EXF
       560/121; 514/530
       163
ARTU
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PATENT KIND DATE

COPYRIGHT 2004 ACS on STN

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9518102 A1
                                            19950706
      CA 123:339522 * WO
OS
                                       Α
                                            20000926
                               6124344
                      US
      CA 133:252211
                               9925358
                                       A1
                                            19990527
                      WO
      CA 131:5147
* CA Indexing for this record included
      26-3 (Biomolecules and Their Synthetic Analogs)
CC
       Section cross-reference(s): 2
      prostaglandin F2a ether prepn ocular hypotensive
ST
      Glaucoma (disease)
IT
        (preparation of prostaglandin derivs. as ocular hypotensives)
                      170753-73-8P
      170753-66-9P
IT
        (preparation of prostaglandin derivs. as ocular hypotensives)
                                                                  170753-67-0P
                                   136198-86-2P
                                                   170753-65-8P
      73726-97-3P
                    79743-27-4P
IT
                                                                     170753-72-7P
                                     170753-70-5P
                                                     170753-71-6P
                      170753-69-2P
      170753-68-1P
                                                                     170753-78-3P
                                     170753-76-1P
                                                     170753-77-2P
                      170753-75-0P
      170753-74-9P
                                                                     170753-83-0P
                                     170753-81-8P
                                                     170753-82-9P
                      170753-80-7P
      170753-79-4P
                                                                     170753-88-5P
                                     170753-86-3P
                                                     170753-87-4P
      170753-84-1P
                      170753-85-2P
                                                     170753-92-1P
                                     170753-91-0P
                      170753-90-9P
      170753-89-6P
                                                                     170753-97-6P
                                                     170753-96-5P
                                     170753-95-4P
                      170753-94-3P
      170753-93-2P
      170753-98-7P
         (preparation of prostaglandin derivs. as ocular hypotensives)
                                                  170753-99-8P
                                  73726-96-2P
                     73726-94-0P
      40834-99-9P
TT
         (preparation of prostaglandin derivs. as ocular hypotensives)
                                     33854-16-9, Prostaglandin F2\alpha
      551-11-1, Prostaglandin F2\alpha
IT
                                                  170754-01-5
                                                                 170754-02-6
                                   170754-00-4
      methyl ester
                      53764-90-2
         (preparation of prostaglandin derivs. as ocular hypotensives)
                     65844-25-9P
                                   65844-26-0P
      63598-54-9P
IT
         (preparation of prostaglandin derivs. as ocular hypotensives)
    170753-89-6P
IT
         (preparation of prostaglandin derivs. as ocular hypotensives)
     170753-89-6 USPATFULL
RN
     Prosta-5,13-dien-1-amide, 9,11-dihydroxy-15-methoxy-,
CN
        (5Z, 9\alpha, 11\alpha, 13E, 15S) - (9CI) (CA INDEX NAME)
```

Absolute stereochemistry.
Double bond geometry as shown.

```
ANSWER 8 OF 14 USPATFULL on STN
L19
       1999:61175 USPATFULL
AN
       Cyclopentane(ENE) heptenoic or heptanoic acids and derivatives thereof
TI
       useful as therapeutic agents
       Burk, Robert M., Laguna Beach, CA, United States
IN
       Allergan Sales, Inc., Irvine, CA, United States (U.S. corporation)
PA
                               19990525
       US 5906989
PΤ
                               19980526 (9)
       US 1998-84805
AΤ
       Division of Ser. No. US 1997-861414, filed on 21 May 1997, now patented,
RLI
       Pat. No. US 5798378 which is a division of Ser. No. US 1996-740883,
       filed on 4 Nov 1996, now patented, Pat. No. US 5681848, issued on 28 Oct
```

170753-97-6P

170753-96-5P

```
1997 which is a division of Ser. No. US 1995-445842, filed on 11 Jul
      1995, now patented, Pat. No. US 5587391, issued on 24 Dec 1996 which is
      a division of Ser. No. US 1993-174535, filed on 28 Dec 1993, now
      patented, Pat. No. US 5545665, issued on 13 Aug 1996
DT
      Utility
      Granted
FS
      Primary Examiner: Lambkin, Deborah C.
EXNAM
      Baran, Robert J., Voet, Martin A., Fisher, Carlos A.
LREP
      Number of Claims: 22
CLMN
       Exemplary Claim: 1
ECL
       4 Drawing Figure(s); 4 Drawing Page(s)
DRWN
LN.CNT 1134
CAS INDEXING IS AVAILABLE FOR THIS PATENT.
       The invention relates to 7-[5-hydroxy-2-(hydroxyhydrocarbyl or
AB
       heteroatom-substituted hydroxy hydrocarbyl)-3-hydroxy-cyclopentyl(enyl)]
       heptanoic or heptenoic acids and derivatives of said acids, wherein one
       or more of said hydroxy groups are replaced by an ether group. The
       compounds of the present invention are potent ocular hypotensives, and
       are particularly suitable for the management of glaucoma.
CAS INDEXING IS AVAILABLE FOR THIS PATENT.
       INCLM: 514/357.000
INCL
       INCLS: 514/277.000; 514/471.000; 514/461.000; 514/530.000; 514/532.000;
              514/561.000; 514/570.000; 514/646.000; 546/329.000; 546/334.000;
              546/339.000; 546/340.000; 546/341.000; 549/491.000; 549/496.000
NCL
       NCLM:
              514/357.000
              514/277.000; 514/461.000; 514/471.000; 514/530.000; 514/532.000;
       NCLS:
              514/561.000; 514/570.000; 514/646.000; 546/329.000; 546/334.000;
              546/339.000; 546/340.000; 546/341.000; 549/491.000; 549/496.000
       [6]
IC
       ICM: A61K031-44
       ICS: A61K031-505; C07D211-70; C07D209-04
       546/329; 546/334; 546/339-341; 514/357; 514/277; 514/471; 514/461;
EXF
       514/530; 514/532; 514/561; 514/570; 514/646; 549/491; 549/496
ARTU
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CHEMICAL ABSTRACTS INDEXING
                               COPYRIGHT 2004 ACS on STN
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                                      KIND
                         PATENT
                    _____ ----
      CA 123:339522 * WO 9518102 A1
                                           19950706
OS
                             6124344 A 20000926
9925358 A1 19990527
      CA 133:252211 US
                    WO
      CA 131:5147
* CA Indexing for this record included
      26-3 (Biomolecules and Their Synthetic Analogs)
CC
       Section cross-reference(s): 2
      prostaglandin F2a ether prepn ocular hypotensive
ST
      Glaucoma (disease)
IT
        (preparation of prostaglandin derivs. as ocular hypotensives)
                     170753-73-8P
      170753-66-9P
IT
        (preparation of prostaglandin derivs. as ocular hypotensives)
                                                 170753-65-8P 170753-67-0P
                    79743-27-4P 136198-86-2P
IT
      73726-97-3P
                                                                  170753-72-7P
                                                   170753-71-6P
                                    170753-70-5P
                     170753-69-2P
      170753-68-1P
                                                                  170753-78-3P
                                                   170753-77-2P
                     170753-75-0P
                                    170753-76-1P
      170753-74-9P
                                                                  170753-83-0P
                                                   170753-82-9P
                     170753-80-7P
                                    170753-81-8P
      170753-79-4P
                                                                  170753-88-5P
                                                   170753-87-4P
                     170753-85-2P
                                    170753-86-3P
      170753-84-1P
                     170753-90-9P
                                    170753-91-0P
                                                   170753-92-1P
      170753-89-6P
```

(preparation of prostaglandin derivs. as ocular hypotensives)

170753-95-4P

170753-94-3P

170753-93-2P 170753-98-7P

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73726-96-2P
                                                  170753-99-8P
                    73726-94-0P
IT
      40834-99-9P
        (preparation of prostaglandin derivs. as ocular hypotensives)
                                    33854-16-9, Prostaglandin F2lpha
      551-11-1, Prostaglandin F2α
IT
                                   170754-00-4 170754-01-5
                                                                170754-02-6
      methyl ester
                     53764-90-2
        (preparation of prostaglandin derivs. as ocular hypotensives)
                                   65844-26-0P
                    65844-25-9P
IT
      63598-54-9P
        (preparation of prostaglandin derivs. as ocular hypotensives)
    170753-89-6P
TT
        (preparation of prostaglandin derivs. as ocular hypotensives)
     170753-89-6 USPATFULL
RN
     Prosta-5,13-dien-1-amide, 9,11-dihydroxy-15-methoxy-,
CN
       (5Z, 9\alpha, 11\alpha, 13E, 15S) - (9CI)
                                   (CA INDEX NAME)
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CAS INDEXING IS AVAILABLE FOR THIS PATENT.

INCLM: 514/438.000

TNCL

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ANSWER 9 OF 14 USPATFULL on STN
L19
       1998:101669 USPATFULL
AN
       Cyclopentane (ene) heptenoic or heptanoic acids and derivatives thereof
TI
       useful as therapeutic agents
       Burk, Robert M., Laguna Beach, CA, United States
IN
       Allergan, Waco, TX, United States (U.S. corporation)
PA
                               19980825
       US 5798378
PΙ
                               19970521 (8)
       US 1997-861414
AΙ
       Division of Ser. No. US 1996-740883, filed on 4 Nov 1996, now patented,
RLI
       Pat. No. US 5681848 which is a division of Ser. No. US 1995-445842,
       filed on 11 Jul 1995, now patented, Pat. No. US 5587391 which is a
       division of Ser. No. US 1993-174535, filed on 28 Dec 1993, now patented,
       Pat. No. US 5545665
       Utility
DT
       Granted
FS
       Primary Examiner: Lambkin, Deborah C.
EXNAM
       Baran, Robert J., Voet, Martin A., Lambert, Howard R.
LREP
       Number of Claims: 8
CLMN
       Exemplary Claim: 1
ECL
       4 Drawing Figure(s); 4 Drawing Page(s)
DRWN
LN.CNT 1018
CAS INDEXING IS AVAILABLE FOR THIS PATENT.
       The invention relates to 7-[5-hydroxy-2-(hydroxyhydrocarbyl or
AB
       heteroatom-substituted hydroxy hydrocarbyl)-3-hydroxycyclopentyl(enyl)]
       heptanoic or heptenoic acids and derivatives of said acids, wherein one
       or more of said hydroxy groups are replaced by an ether group. The
       compounds of the present invention are potent ocular hypotensives, and
       are particularly suitable for the management of glaucoma.
```

```
INCLS: 549/075.000; 549/076.000; 549/077.000; 549/078.000; 549/079.000
NCL
       NCLM:
              514/438.000
             549/075.000; 549/076.000; 549/077.000; 549/078.000; 549/079.000
       NCLS:
       [6]
IC
       ICM: A61K031-38
       ICS: C07D333-12; C07D333-16; C07D333-24
       574/438; 549/75; 549/76; 549/77; 549/78; 549/79; 549/74
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                                      KIND
                                              DATE
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OS
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                                       A1
      CA 133:252211
                      US
                             6124344
                                       Α
                                            20000926
                              9925358 A1
                                            19990527
      CA 131:5147
                      WO
* CA Indexing for this record included
      26-3 (Biomolecules and Their Synthetic Analogs)
CC
       Section cross-reference(s): 2
      prostaglandin F2a ether prepn ocular hypotensive
ST
      Glaucoma (disease)
IT
        (preparation of prostaglandin derivs. as ocular hypotensives)
IT
      170753-66-9P
                     170753-73-8P
```

(preparation of prostaglandin derivs. as ocular hypotensives)

170753-67-0P 170753-65-8P 79743-27-4P 136198-86-2P IT 73726-97-3P 170753-71-6P 170753-72-7P 170753-70-5P 170753-68-1P 170753-69-2P 170753-77-2P 170753-78-3P 170753-76-1P 170753-74-9P 170753-75-0P 170753-83-0P 170753-82-9P 170753-80-7P 170753-81-8P 170753-79-4P 170753-88-5P 170753-87-4P 170753-85-2P 170753-86-3P 170753-84-1P 170753-92-1P 170753-91-0P 170753-89-6P 170753-90-9P 170753-96-5P 170753-95-4P 170753-97-6P 170753-93-2P 170753-94-3P 170753-98-7P

(preparation of prostaglandin derivs. as ocular hypotensives)

73726-94-0P 73726-96-2P 170753-99-8P IT 40834-99-9P

(preparation of prostaglandin derivs. as ocular hypotensives)

551-11-1, Prostaglandin F2 $\alpha$  33854-16-9, Prostaglandin F2 $\alpha$ 170754-02-6 170754-01-5 170754-00-4 53764-90-2 methyl ester (preparation of prostaglandin derivs. as ocular hypotensives)

65844-26-0P 65844-25-9P 63598-54-9P

(preparation of prostaglandin derivs. as ocular hypotensives)

170753-89-6P IT

TT

IT

(preparation of prostaglandin derivs. as ocular hypotensives)

170753-89-6 USPATFULL RN

Prosta-5,13-dien-1-amide, 9,11-dihydroxy-15-methoxy-, CN $(5Z, 9\alpha, 11\alpha, 13E, 15S) - (9CI)$  (CA INDEX NAME)

```
L19 ANSWER 10 OF 14 USPATFULL on STN
      97:99299 USPATFULL
AN
      Cyclopentane (ene) heptenoic or heptanoic acids and derivatives thereof
ΤI
      useful as therapeutic agents
       Burk, Robert M., Laguna Beach, CA, United States
TN
      Allergan, Waco, TX, United States (U.S. corporation)
PA
                              19971028
      US 5681848
PΙ
                              19961104 (8)
      US 1996-740883
AΙ
      Division of Ser. No. US 1995-445842, filed on 11 Jul 1995, now patented,
RLI
       Pat. No. US 5587391 which is a division of Ser. No. US 1993-174535,
       filed on 28 Dec 1993, now patented, Pat. No. US 5545665, issued on 13
       Aug 1996
       Utility
DT
       Granted
FS
      Primary Examiner: Dees, Jose G.; Assistant Examiner: Cebulak, Mary C.
EXNAM
       Baran, Robert J., Voet, Martin A., Lambert, Howard R.
LREP
       Number of Claims: 8
CLMN
       Exemplary Claim: 1
ECL
       4 Drawing Figure(s); 4 Drawing Page(s)
DRWN
LN.CNT 1007
CAS INDEXING IS AVAILABLE FOR THIS PATENT.
       The invention relates to 7-[5-hydroxy-2-(hydroxyhydrocarbyl or
       heteroatom-substituted hydroxy hydrocarbyl)-3-
       hydroxycyclopentyl(enyl)]heptanoic or heptenoic acids and derivatives of
       said adds, wherein one or more of said hydroxy groups are replaced by an
       ether group. The compounds of the present invention are potent ocular
       hypotensives, and are particularly suitable for the management of
       glaucoma.
CAS INDEXING IS AVAILABLE FOR THIS PATENT.
       INCLM: 514/471.000
INCL
       INCLS: 514/912.000; 549/498.000
       NCLM: 514/471.000
NCL
       NCLS: 514/912.000; 549/498.000
IC
       [6]
       ICM: A61K031-34
       ICS: C07D307-36
       514/471; 514/912; 549/498
EXF
       129
ARTU
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CHEMICAL ABSTRACTS INDEXING
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                                     KIND
                                            DATE
                         PATENT
                    ______
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OS
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                                          20000926
      CA 133:252211 US
                     WO
                             9925358 A1 19990527
      CA 131:5147
* CA Indexing for this record included
      26-3 (Biomolecules and Their Synthetic Analogs)
       Section cross-reference(s): 2
      prostaglandin F2a ether prepn ocular hypotensive
st
      Glaucoma (disease)
IT
        (preparation of prostaglandin derivs. as ocular hypotensives)
      170753-66-9P
                     170753-73-8P
TТ
        (preparation of prostaglandin derivs. as ocular hypotensives)
                                  136198-86-2P 170753-65-8P 170753-67-0P
      73726-97-3P 79743-27-4P
TT
                                    170753-70-5P 170753-71-6P
                                                                 170753-72-7P
                    170753-69-2P
      170753-68-1P
```

170753-76-1P

170753-74-9P 170753-75-0P

170753-77-2P

170753-78-3P

```
170753-82-9P
                                                                     170753-83-0P
                     170753-80-7P
                                     170753-81-8P
      170753-79-4P
                                     170753-86-3P
                                                     170753-87-4P
                                                                     170753-88-5P
                     170753-85-2P
      170753-84-1P
                                     170753-91-0P
                                                     170753-92-1P
                     170753-90-9P
      170753-89-6P
                                                     170753-96-5P
                                                                     170753-97-6P
                     170753-94-3P
                                     170753-95-4P
      170753-93-2P
      170753-98-7P
        (preparation of prostaglandin derivs. as ocular hypotensives)
                                   73726-96-2P
                                                 170753-99-8P
                    73726-94-0P
IT
      40834-99-9P
        (preparation of prostaglandin derivs. as ocular hypotensives)
      551-11-1, Prostaglandin F2\alpha
                                     33854-16-9, Prostaglandin F2\alpha
ΙT
                                   170754-00-4
                                                  170754-01-5
                                                                 170754-02-6
                     53764-90-2
      methyl ester
        (preparation of prostaglandin derivs. as ocular hypotensives)
                    65844-25-9P
                                   65844-26-0P
IT
      63598-54-9P
        (preparation of prostaglandin derivs. as ocular hypotensives)
    170753-89-6P
TT
        (preparation of prostaglandin derivs. as ocular hypotensives)
RN
     170753-89-6 USPATFULL
     Prosta-5,13-dien-1-amide, 9,11-dihydroxy-15-methoxy-,
CN
       (5Z, 9\alpha, 11\alpha, 13E, 15S) - (9CI) (CA INDEX NAME)
```

HO 
$$\frac{Z}{(CH_2)_3}$$
  $NH_2$ 
 $R$   $R$   $E$   $OMe$   $O$ 
 $(CH_2)_4$   $Me$ 

```
ANSWER 11 OF 14 USPATFULL on STN
L19
       96:118603 USPATFULL
AN
       Cyclopentane (ene) heptenoic or heptanoic acids and derivatives thereof
TI
       useful as therapeutic agents
       Burk, Robert M., Laguna Beach, CA, United States
IN
       Allergan, Waco, TX, United States (U.S. corporation)
PA
                               19961224
       US 5587391
PΙ
                               19950711 (8)
       US 1995-445842
AΙ
       Division of Ser. No. US 1993-174535, filed on 28 Dec 1993, now patented,
RLI
       Pat. No. US 5545665
       Utility
דת
       Granted
FS
       Primary Examiner: Raymond, Richard L.; Assistant Examiner: Lambkin,
EXNAM
       Baran, Robert J., Voet, Martin A., Lambert, Howard R.
LREP
       Number of Claims: 9
CLMN
       Exemplary Claim: 1
ECL
       4 Drawing Figure(s); 4 Drawing Page(s)
DRWN
LN.CNT 999
CAS INDEXING IS AVAILABLE FOR THIS PATENT.
       The invention relates to 7-[5-hydroxy-2-(hydroxyhydrocarbyl or
AB
       heteroatom-substituted hydroxy hydrocarbyl)-3-hydroxycyclopentyl(enyl)]
       heptanoic or heptenoic acids and derivatives of said acids, wherein one
       or more of said hydroxy groups are replaced by an ether group. The
       compounds of the present invention are potent ocular hypotensives, and
```

are particularly suitable for the management of glaucoma.

```
CAS INDEXING IS AVAILABLE FOR THIS PATENT.
INCL
      INCLM: 514/357.000
      INCLS: 546/337.000
      NCLM: 514/357.000
NCL
      NCLS: 546/337.000
IC
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      ICM: A61K031-44
      ICS: C07D213-56
      546/290; 546/304; 546/312; 546/311; 546/326; 546/345; 546/340; 546/341;
EXF
      546/344; 546/337; 514/345; 514/347; 514/352; 514/354; 514/357
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      129
CHEMICAL ABSTRACTS INDEXING
                              COPYRIGHT 2004 ACS on STN
                        PATENT
                                     KIND
     CA 123:339522 * WO 9518102 A1 19950706
OS
      CA 133:252211 US
                             6124344 A
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                    WO
     CA 131:5147
                             9925358 A1 19990527
* CA Indexing for this record included
      26-3 (Biomolecules and Their Synthetic Analogs)
CC
       Section cross-reference(s): 2
      prostaglandin F2a ether prepn ocular hypotensive
ST
     Glaucoma (disease)
IT
        (preparation of prostaglandin derivs. as ocular hypotensives)
      170753-66-9P
                    170753-73-8P
TT
        (preparation of prostaglandin derivs. as ocular hypotensives)
      73726-97-3P 79743-27-4P 136198-86-2P 170753-65-8P 170753-67-0P
IT
                                                               170753-72-7P
      170753-68-1P 170753-69-2P 170753-70-5P 170753-71-6P
                                                                170753-78-3P
      170753-74-9P 170753-75-0P 170753-76-1P
                                                  170753-77-2P
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                                                                170753-97-6P
      170753-93-2P 170753-94-3P 170753-95-4P
                                                  170753-96-5P
      170753-98-7P
        (preparation of prostaglandin derivs. as ocular hypotensives)
                   73726-94-0P 73726-96-2P 170753-99-8P
IT
      40834-99-9P
        (preparation of prostaglandin derivs. as ocular hypotensives)
      551-11-1, Prostaglandin F2\alpha 33854-16-9, Prostaglandin F2\alpha
TT
                    53764-90-2 170754-00-4 170754-01-5
                                                             170754-02-6
      methyl ester
        (preparation of prostaglandin derivs. as ocular hypotensives)
      63598-54-9P
                                65844-26-0P
IT
                   65844-25-9P
        (preparation of prostaglandin derivs. as ocular hypotensives)
    170753-89-6P
TT
        (preparation of prostaglandin derivs. as ocular hypotensives)
     170753-89-6 USPATFULL
RN
     Prosta-5,13-dien-1-amide, 9,11-dihydroxy-15-methoxy-,
CN
       (5Z, 9\alpha, 11\alpha, 13E, 15S) - (9CI) (CA INDEX NAME)
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ANSWER 12 OF 14 USPATFULL on STN
L19
       96:72913 USPATFULL
AN
       Cyclopentane(ene) heptenoic or heptanoic acids and derivatives thereof
TI
       useful as therapeutic agents
       Burk, Robert M., Laguna Beach, CA, United States
IN
       Allergan, Waco, TX, United States (U.S. corporation)
PA
                               19960813
       US 5545665
PΙ
                               19931228 (8)
       US 1993-174535
ΑI
DT
       Utility
       Granted
FS
       Primary Examiner: Raymond, Richard L.; Assistant Examiner: Lambkin,
EXNAM
       Deborah
       Baran, Robert J., Voet, Martin A., Lambert, Howard R.
LREP
       Number of Claims: 29
CLMN
       Exemplary Claim: 1
ECL
       4 Drawing Figure(s); 4 Drawing Page(s)
DRWN
LN.CNT 1164
CAS INDEXING IS AVAILABLE FOR THIS PATENT.
       The invention relates to 7-[5-hydroxy-2-(hydroxyhydrocarbyl or
AB
       heteroatom-substituted hydroxy hydrocarbyl)-3-hydroxycyclopentyl(enyl)]
       heptanoic or heptenoic acids and derivatives of said acids, wherein one
       or more of said hydroxy groups are replaced by an ether group. The
       compounds of the present invention are potent ocular hypotensives, and
       are particularly suitable for the management of glaucoma.
CAS INDEXING IS AVAILABLE FOR THIS PATENT.
       INCLM: 514/530.000
INCL
       INCLS: 514/573.000; 514/613.000; 514/659.000; 514/729.000; 560/121.000;
              562/503.000; 562/504.000; 562/510.000; 564/189.000; 564/453.000;
               564/454.000; 568/838.000
              514/530.000
NCL
       NCLS: 514/573.000; 514/613.000; 514/659.000; 514/729.000; 560/121.000;
              562/503.000; 562/504.000; 562/510.000; 564/189.000; 564/453.000;
               564/454.000; 568/838.000
        [6]
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        ICM: A61K031-25
        ICS: A61K031-557; C07C405-00; C07C233-00
        562/503; 562/504; 562/510; 514/530; 514/659; 514/573; 514/613; 514/729;
 EXF
        560/121; 568/838; 564/189; 564/453; 564/454
 ARTU
        129
                                COPYRIGHT 2004 ACS on STN
 CHEMICAL ABSTRACTS INDEXING
                                       KIND DATE
                          PATENT
                              9518102 A1 19950706
       CA 123:339522 * WO
 OS
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20000926
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      CA 133:252211
                      US
                               9925358 A1 19990527
                      WO
      CA 131:5147
* CA Indexing for this record included
      26-3 (Biomolecules and Their Synthetic Analogs)
       Section cross-reference(s): 2
      prostaglandin F2a ether prepn ocular hypotensive
ST
      Glaucoma (disease)
IT
        (preparation of prostaglandin derivs. as ocular hypotensives)
      170753-66-9P
                     170753-73-8P
IT
        (preparation of prostaglandin derivs. as ocular hypotensives)
                                                                  170753-67-0P
                                   136198-86-2P
                                                  170753-65-8P
      73726-97-3P
                    79743-27-4P
IT
                                     170753-70-5P
                                                     170753-71-6P
                                                                    170753-72-7P
                     170753-69-2P
      170753-68-1P
                                                                    170753-78-3P
                                                     170753-77-2P
                                     170753-76-1P
                     170753-75-0P
      170753-74-9P
                                                                    170753-83-0P
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                                                     170753-82-9P
      170753-79-4P
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                                                                     170753-88-5P
                                     170753-86-3P
                                                     170753-87-4P
                     170753-85-2P
      170753-84-1P
                                                     170753-92-1P
                                     170753-91-0P
                     170753-90-9P
      170753-89-6P
                                                                     170753-97-6P
                                                     170753-96-5P
                                     170753-95-4P
                     170753-94-3P
      170753-93-2P
      170753-98-7P
        (preparation of prostaglandin derivs. as ocular hypotensives)
                                                  170753-99-8P
                                   73726-96-2P
                     73726-94-0P
      40834-99-9P
IT
        (preparation of prostaglandin derivs. as ocular hypotensives)
                                     33854-16-9, Prostaglandin F2\alpha
      551-11-1, Prostaglandin F2lpha
IT
                                                                170754-02-6
                                   170754-00-4
                                                  170754-01-5
                      53764-90-2
      methyl ester
         (preparation of prostaglandin derivs. as ocular hypotensives)
                                   65844-26-0P
                     65844-25-9P
      63598-54-9P
IT
         (preparation of prostaglandin derivs. as ocular hypotensives)
    170753-89-6P
IT
         (preparation of prostaglandin derivs. as ocular hypotensives)
     170753-89-6 USPATFULL
RN
     Prosta-5,13-dien-1-amide, 9,11-dihydroxy-15-methoxy-,
CN
        (5Z, 9\alpha, 11\alpha, 13E, 15S) - (9CI) (CA INDEX NAME)
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ANSWER 13 OF 14 USPAT2 on STN
L19
       2002:259478 USPAT2
AN
       Cyclopentane (ENE) heptenoic or heptanoic acids and derivatives thereof
ΤI
       useful as therapeutic agents
       Burk, Robert M., Laguna Beach, CA, United States
IN
       Allergan, Inc., Irvine, CA, United States (U.S. corporation)
PA
                               20040406
                          B2
       US 6716876
PI
                               20020228 (10)
       US 2002-87867
ΑT
       Continuation of Ser. No. US 2001-919318, filed on 31 Jul 2001
RLI
       Continuation of Ser. No. US 1999-448082, filed on 23 Nov 1999, now
       patented, Pat. No. US 6303658 Continuation of Ser. No. US 1999-225034,
       filed on 4 Jan 1999, now patented, Pat. No. US 5990138, issued on 23 Nov
```

1999 Division of Ser. No. US 1998-84805, filed on 26 May 1998, now patented, Pat. No. US 5906989, issued on 25 May 1999 Division of Ser. No. US 1997-861414, filed on 21 May 1997, now patented, Pat. No. US 5798378, issued on 25 Aug 1998 Division of Ser. No. US 1996-740883, filed on 4 Nov 1996, now patented, Pat. No. US 5681848, issued on 28 Oct 1997 Division of Ser. No. US 1995-445842, filed on 11 Jul 1995, now patented, Pat. No. US 5587391, issued on 4 Dec 1996 Division of Ser. No. US 1993-174535, filed on 28 Dec 1993, now patented, Pat. No. US 5545665, issued on 13 Aug 1996

DTUtility GRANTED FS

Primary Examiner: Gorsth, Robert EXNAM

Baran, Robert J., Voet, Martin A., Fisher, Carlos A. LREP

Number of Claims: 1 CLMN Exemplary Claim: 1 ECL

4 Drawing Figure(s); 4 Drawing Page(s) DRWN

LN.CNT 978

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

The invention relates to 7-[5-hydroxy-2-(hydroxyhydrocarbyl or AΒ heteroatom-substituted hydroxy hydrocarbyl)-3-hydroxy-cyclopentyl(enyl)] heptanoic or heptenoic acids and derivatives of said acids, wherein one or more of said hydroxy groups are replaced by an ether group. The compounds of the present invention are potent ocular hypotensives, and are particularly suitable for the management of glaucoma.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

INCLM: 514/530.000 INCL

INCLS: 514/573.000; 514/546.000; 514/568.000; 514/613.000; 514/715.000

514/530.000 NCL NCLM:

514/546.000; 514/568.000; 514/573.000; 514/613.000; 514/715.000 NCLS:

DATE

IC [7]

ICM: A61K031-5575

514/530; 514/573; 514/346; 514/568; 514/613; 514/715 EXF

ARTU

## COPYRIGHT 2004 ACS on STN CHEMICAL ABSTRACTS INDEXING

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                                     KIND
                            9518102 Al 19950706
     CA 123:339522 * WO
OS
                             6124344 A
                                          20000926
     CA 133:252211
                     US
                             9925358 A1 19990527
     CA 131:5147 WO
* CA Indexing for this record included
      26-3 (Biomolecules and Their Synthetic Analogs)
CC
      Section cross-reference(s): 2
     prostaglandin F2a ether prepn ocular hypotensive
ST
IT
     Glaucoma (disease)
        (preparation of prostaglandin derivs. as ocular hypotensives)
                    170753-73-8P
     170753-66-9P
IT
        (preparation of prostaglandin derivs. as ocular hypotensives)
                  79743-27-4P 136198-86-2P 170753-65-8P
                                                              170753-67-0P
      73726-97-3P
IT
                                                                170753-72-7P
                                   170753-70-5P
                                                  170753-71-6P
                    170753-69-2P
      170753-68-1P
                                                  170753-77-2P
                                                                 170753-78-3P
                                   170753-76-1P
     170753-74-9P
                    170753-75-0P
                                   170753-81-8P
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      170753-84-1P
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                    170753-90-9P
      170753-89-6P
                                                                 170753-97-6P
                                   170753-95-4P
                                                  170753-96-5P
                    170753-94-3P
      170753-93-2P
      170753-98-7P
        (preparation of prostaglandin derivs. as ocular hypotensives)
                  73726-94-0P 73726-96-2P 170753-99-8P
IT
      40834-99-9P
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```
(preparation of prostaglandin derivs. as ocular hypotensives)
      551-11-1, Prostaglandin F2\alpha 33854-16-9, Prostaglandin F2\alpha
IT
                                                                 170754-02-6
                                                  170754-01-5
                                   170754-00-4
      methyl ester
                      53764-90-2
        (preparation of prostaglandin derivs. as ocular hypotensives)
                                   65844-26-0P
                    65844-25-9P
IT
      63598-54-9P
        (preparation of prostaglandin derivs. as ocular hypotensives)
    170753-89-6P
IT
        (preparation of prostaglandin derivs. as ocular hypotensives)
     170753-89-6 USPAT2
RN
     Prosta-5,13-dien-1-amide, 9,11-dihydroxy-15-methoxy-,
CN
       (5Z, 9\alpha, 11\alpha, 13E, 15S) - (9CI) (CA INDEX NAME)
```

HO 
$$\frac{Z}{(CH_2)_3}$$
  $\frac{NH_2}{NH_2}$  HO  $\frac{R}{R}$   $\frac{E}{(CH_2)_4}$   $\frac{NH_2}{Me}$ 

```
ANSWER 14 OF 14 USPAT2 on STN
L19
       2002:4171 USPAT2
AN
      Cyclopentane (ENE) heptenoic or heptenoic acids and derivatives thereof
ΤI
       useful as therapeutic agents
       Burk, Robert M., Laguna Beach, CA, United States
IN
      Allergan Sales, Inc., Irvine, CA, United States (U.S. corporation)
PA
                               20020702
                          В2
       US 6414022
PΙ
                               20010731 (9)
       US 2001-919318
AI
       Continuation of Ser. No. US 1999-448082, filed on 23 Nov 1999, now
RLI
       patented, Pat. No. US 6303658 Continuation of Ser. No. US 1999-225034,
       filed on 4 Jan 1999, now patented, Pat. No. US 5990138, issued on 23 Nov
       1999 Division of Ser. No. US 1998-84805, filed on 26 May 1998, now
       patented, Pat. No. US 5906989, issued on 25 May 1999 Division of Ser.
       No. US 1997-861414, filed on 21 May 1997, now patented, Pat. No. US
       5798378, issued on 25 Aug 1998 Division of Ser. No. US 1996-740883,
       filed on 4 Nov 1996, now patented, Pat. No. US 5681848, issued on 28 Oct
       1997 Division of Ser. No. US 1995-445842, filed on 11 Jul 1995, now
       patented, Pat. No. US 5587391, issued on 4 Dec 1996 Division of Ser. No.
       US 1993-174535, filed on 28 Dec 1993, now patented, Pat. No. US 5545665,
       issued on 13 Aug 1996
DT
       Utility
       GRANTED
FS
       Primary Examiner: Gerstl, Robert
EXNAM
       Baran, Robert J., Voet, Martin A., Fisher, Carlos A.
LREP
       Number of Claims: 7
CLMN
       Exemplary Claim: 1
ECL
       4 Drawing Figure(s); 4 Drawing Page(s)
DRWN
LN.CNT 1081
CAS INDEXING IS AVAILABLE FOR THIS PATENT.
       The invention relates to 7-[5-hydroxy-2-(hydroxyhydrocarbyl or
       heteroatom-substituted hydroxy hydrocarbyl)-3-hydroxy-cyclopentyl(enyl)]
       heptanoic or heptenoic acids and derivatives of said acids, wherein one
```

or more of said hydroxy groups are replaced by an ether group. The compounds of the present invention are potent ocular hypotensives, and are particularly suitable for the management of glaucoma.

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INCLM: 514/530.000
INCL
      INCLS: 514/573.000; 514/546.000; 514/568.000; 514/613.000; 514/715.000
      NCLM: 514/530.000
NCL
      NCLS: 514/546.000; 514/568.000; 514/573.000; 514/613.000; 514/715.000
IC
       [7]
      ICM: A61K031-5575
      514/530; 514/573; 514/715; 514/568; 514/613; 514/546
EXF
ARTU
CHEMICAL ABSTRACTS INDEXING
                              COPYRIGHT 2004 ACS on STN
                                     KIND
                         PATENT
                    _______
      CA 123:339522 * WO
                             9518102 A1 19950706
OS
      CA 133:252211 US
                             6124344 A
                                           20000926
      CA 131:5147 WO
                             9925358 A1 19990527
* CA Indexing for this record included
      26-3 (Biomolecules and Their Synthetic Analogs)
CC
       Section cross-reference(s): 2
      prostaglandin F2a ether prepn ocular hypotensive
ST
TT
      Glaucoma (disease)
        (preparation of prostaglandin derivs. as ocular hypotensives)
                     170753-73-8P
      170753-66-9P
IT
        (preparation of prostaglandin derivs. as ocular hypotensives)
                   79743-27-4P 136198-86-2P 170753-65-8P 170753-67-0P
      73726-97-3P
TT
                                                                  170753-72-7P
                                    170753-70-5P
                                                   170753-71-6P
      170753-68-1P
                    170753-69-2P
                                                                  170753-78-3P
                                    170753-76-1P
                                                   170753-77-2P
      170753-74-9P
                     170753-75-0P
                                                                  170753-83-0P
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      170753-79-4P
                                    170753-86-3P
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      170753-84-1P
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                                                   170753-96-5P 170753-97-6P
      170753-93-2P
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                                    170753-95-4P
      170753-98-7P
        (preparation of prostaglandin derivs. as ocular hypotensives)
                   73726-94-0P 73726-96-2P
                                              170753-99-8P
      40834-99-9P
IT
        (preparation of prostaglandin derivs. as ocular hypotensives)
      551-11-1, Prostaglandin F2\alpha 33854-16-9, Prostaglandin F2\alpha
IT
                                                              170754-02-6
                     53764-90-2 170754-00-4
                                                170754-01-5
      methyl ester
        (preparation of prostaglandin derivs. as ocular hypotensives)
                                  65844-26-0P
                    65844-25-9P
      63598-54-9P
IT
        (preparation of prostaglandin derivs. as ocular hypotensives)
    170753-89-6P
IT
        (preparation of prostaglandin derivs. as ocular hypotensives)
     170753-89-6 USPAT2
RN
     Prosta-5,13-dien-1-amide, 9,11-dihydroxy-15-methoxy-,
CN
       (5Z, 9\alpha, 11\alpha, 13E, 15S) - (9CI) (CA INDEX NAME)
```

Absolute stereochemistry.

Double bond geometry as shown.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

HO 
$$\frac{Z}{(CH_2)_3}$$
  $\frac{NH_2}{NH_2}$  HO  $\frac{R}{R}$   $\frac{E}{(CH_2)_4}$   $\frac{NH_2}{Me}$ 

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